

A' disease in the individual as compared with an individual having one or more of a guanine at nucleotide position 610, a cytosine at nucleotide position 514, a guanine at nucleotide position 218, a thymine at nucleotide position 425, an adenine at nucleotide position 197, an adenine at nucleotide position 112, a guanine at nucleotide position 233, a thymine at nucleotide position 608, a guanine at nucleotide position 143, a guanine at nucleotide position 317, and a cytosine at nucleotide position 316, respectively.

11. (New) A method according to Claim 4, wherein the inflammatory bowel disease is Crohn's disease.
 12. (New) A method according to Claim 4, wherein the individual is an individual at risk for development of Crohn's disease.
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REMARKS

Amendments to the Specification

✓ The Substitute "Sequence Listing" is being submitted to include 300 sequences from Table 3 that were inadvertently omitted from the "Sequence Listing" as previously filed. Support for the Substitute Sequence listing can be found in Table 3; no new matter has been added.

The Substitute Specification is being submitted to include Tables 3 and 5 as amended. Tables 3 and 5 are being amended to identify the sequence disclosures with SEQ ID NOS as required by MPEP §2422. Additionally, the heading row from the first page of Table 3 has been added to each subsequent page of the table. No new matter has been added.

Claim Amendments

Support for new Claims 7-12 can be found throughout the Specification, for example, at page 5, lines 15-25 and at page 31, lines 15-31; no new matter has been added.

Restriction of Claims 1-6 as Originally Filed Under 35 §U.S.C. 121

The Examiner states that restriction to one of the inventions from Table 3, as recited in Claims 1-6 as originally filed, is required.

Claims 1-6 as originally filed have been cancelled, thus rendering the issue of the restriction for these claims moot.

New Claims 7-12 are presented, which recite a Markush group of 12 single nucleotide polymorphisms (SNPs) which show strong linkage disequilibrium with inflammatory bowel disease. These 12 SNPs form a haplotype, which is a segment of a chromosome(s) that contain genetic variations inherited together as a set or a block. A haplotype can be used to decipher the genetic differences that result in phenotypic differences between individuals. For example, identification of the haplotype described in the subject application which is associated with inflammatory bowel diseases will aid in the diagnosis and treatment of these diseases.

MPEP §803.02 states that if members of a Markush group are sufficiently few in number or so closely related that a search and examination of the entire claim can be made without serious burden, the Examiner must examine all the members of the Markush group in the claim on the merits, even though they are directed to independent and distinct inventions. Since the decisions in *In re Weber*, 198 USPQ 328 (CCPA 1978) and *In re Haas*, 198 USPQ 334 (CCPA 1978), it is improper for the Patent Office to refuse to examine that which Applicants regard as their invention, unless the subject matter in a claim lacks unity of invention. Unity of invention exists where compounds included within a Markush group (1) share a common utility, and (2) share a substantial structural feature disclosed as being essential to that utility. As the SNPs presented in these claims satisfy the requirements of unity of invention, as discussed above, Applicants believe that these claims should be examined in their entirety.

In applications containing Markush-type claims which include independent and distinct inventions, the Examiner may require a provisional election of a single species prior to examination on the merits. The provisional election will be given effect in the event that the Markush-type claim should be found not allowable. Following election, the Markush-type claim will be examined fully with respect to the elected species and further to the extent necessary to

determine patentability. If the Markush-type claim is not allowable over the prior art, examination will be limited to the Markush-type claim and claims to the elected species, with claims drawn to species patentably distinct from the elected species held withdrawn from further consideration (See MPEP §803.02).

In that event, Applicants provisionally elect the SNP located at nucleotide position 218 relative to the 5'-most start codon in SEQ ID NO: 1127 (which can be found in Table 5, on page 108).

CONCLUSION

In view of the above amendments and remarks, it is believed that all claims are in condition for allowance, and it is respectfully requested that the application be passed to issue. If the Examiner feels that a telephone conference would expedite prosecution of this case, the Examiner is invited to call the undersigned at (978) 341-0036.

Respectfully submitted,

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6/6/02

MARKED UP VERSION OF AMENDMENTSSpecification Amendments Under 37 C.F.R. § 1.121(b)(1)(iii)

Please amend the Specification by replacing the Specification as filed (pages 1-486) with the enclosed Substitute Specification (pages 1-486). A marked up version of the Substitute Specification showing all the changes relative to the Specification as filed is attached as Exhibit A; deleted matter is indicated with brackets, and added matter is underlined.

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PATENT APPLICATION
Docket No.: 2825.1025-002

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Attorney's Docket No.: 2825.1025-002

Crohn's Disease-Related Polymorphisms
~~IBD-RELATED POLYMORPHISMS~~

RELATED APPLICATIONS

This application claims the benefit of U.S. Provisional Application Serial No. 60/170,257, filed December 10, 1999, and U.S. Provisional Application Serial No. 60/196,046, filed April 10, 2000, the entire teachings of both of which are incorporated herein by reference in their entirety.

BACKGROUND OF THE INVENTION

The genomes of all organisms undergo spontaneous mutation in the course of their continuing evolution, generating variant forms of progenitor nucleic acid sequences (Gusella, *Ann. Rev. Biochem.* 55, 831-854 (1986)). The variant form may confer an evolutionary advantage or disadvantage relative to a progenitor form, or may be neutral. In some instances, a variant form confers a lethal disadvantage and is not transmitted to subsequent generations of the organism. In other instances, a variant form confers an evolutionary advantage to the species and is eventually incorporated into the DNA of many or most members of the species and effectively becomes the

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progenitor form. In many instances, both progenitor and variant form(s) survive and co-exist in a species population. The coexistence of multiple forms of a sequence gives rise to polymorphisms.

Several different types of polymorphism have been reported. A restriction
5 fragment length polymorphism (RFLP) is a variation in DNA sequence that alters the
length of a restriction fragment (Botstein *et al.*, *Am. J. Hum. Genet.* 32, 314-331
(1980)). The restriction fragment length polymorphism may create or delete a
restriction site, thus changing the length of the restriction fragment. RFLPs have been
widely used in human and animal genetic analyses (see WO 90/13668; W090/11369;
10 Donis-Keller, *Cell* 51, 319-337 (1987); Lander *et al.*, *Genetics* 121, 85-99 (1989)).
When a heritable trait can be linked to a particular RFLP, the presence of the RFLP in
an individual can be used to predict the likelihood that the animal will also exhibit the
trait.

Other polymorphisms take the form of short tandem repeats (STRs) that include
15 tandem di-, tri- and tetra-nucleotide repeated motifs. These tandem repeats are also
referred to as variable number tandem repeat (VNTR) polymorphisms. VNTRs have
been used in identity and paternity analysis (US 5,075,217; Armour *et al.*, *FEBS Lett.*
307, 113-115 (1992); Horn *et al.*, W0 91/14003; Jeffreys, EP 370,719), and in a large
number of genetic mapping studies.

20 Other polymorphisms take the form of single nucleotide variations between
individuals of the same species. Such polymorphisms are far more frequent than
RFLPs, STRs and VNTRs. Some single nucleotide polymorphisms (SNP) occur in
protein-coding nucleic acid sequences (coding sequence SNP (cSNP)), in which case,
one of the polymorphic forms may give rise to the expression of a defective or otherwise
25 variant protein and, potentially, a genetic disease. Examples of genes in which
polymorphisms within coding sequences give rise to genetic disease include β -globin
(sickle cell anemia), apoE4 (Alzheimer's Disease), Factor V Leiden (thrombosis), and
CFTR (cystic fibrosis). cSNPs can alter the codon sequence of the gene and therefore
specify an alternative amino acid. Such changes are called "missense" when another

amino acid is substituted, and "nonsense" when the alternative codon specifies a stop signal in protein translation. When the cSNP does not alter the amino acid specified the cSNP is called "silent". Other single nucleotide polymorphisms occur in noncoding regions. Some of these polymorphisms may also result in defective protein expression
5 (e.g., as a result of defective splicing). Other single nucleotide polymorphisms have no phenotypic effects.

Single nucleotide polymorphisms can be used in the same manner as RFLPs and VNTRs, but offer several advantages. Single nucleotide polymorphisms occur with greater frequency and are spaced more uniformly throughout the genome than other
10 forms of polymorphism. The greater frequency and uniformity of single nucleotide polymorphisms means that there is a greater probability that such a polymorphism will be found in close proximity to a genetic locus of interest than would be the case for other polymorphisms. The different forms of characterized single nucleotide polymorphisms are often easier to distinguish than other types of polymorphism (e.g.,
15 by use of assays employing allele-specific hybridization probes or primers).

Only a small percentage of the total repository of polymorphisms in humans and other organisms has been identified. The limited number of polymorphisms identified to date is due to the large amount of work required for their detection by conventional methods. For example, a conventional approach to identifying polymorphisms might be
20 to sequence the same stretch of DNA in a population of individuals by dideoxy sequencing. In this type of approach, the amount of work increases in proportion to both the length of sequence and the number of individuals in a population and becomes impractical for large stretches of DNA or large numbers of persons.

SUMMARY OF THE INVENTION

25 Work described herein pertains to the identification of polymorphisms which are associated with inflammatory bowel diseases (IBD), and particularly those within a single risk haplotype, by resequencing large numbers of genes and gene fragments in a large number of individuals. Various genes from a number of individuals have been

resequenced as described herein, and SNPs in these genes have been discovered (see Table 3). Some of these SNPs are cSNPs which specify a different amino acid sequence, some of the SNPs are silent cSNPs and some of these cSNPs specify a stop signal in protein translation. Some of the identified SNPs were located in non-coding regions.

With the goal of identifying IBD susceptibility genes, a genomewide scan was undertaken in 163 pedigrees, and three regions of suggestive linkage (3, 5q31-33, 6p) and one of significant linkage to 19p13 (LOD = 4.6) were identified. Higher density mapping in the suggestive 5q31-33 region revealed a CD susceptibility locus of genome-wide significance (LOD = 3.9). Importantly, the 5q31-p33 localizes to the major immunoregulatory cytokine gene cluster and the 19p13 locus to a region containing numerous genes encoding cytokine/chemokine receptors and other inflammatory-associated molecules that could have a direct role in disease susceptibility.

In order to pursue the evidence of linkage to chromosome 5, a systematic linkage disequilibrium (LD) approach was adopted. The approach that was used in the first stage of LD mapping was to genotype all known microsatellite markers in the 18 cM between D5S1435 and D5S1480, as these two markers delimit a region of a 2 LOD drop on either side of the linkage peak centered at marker D5S2497. All alleles for each marker were examined for evidence of excess transmission from heterozygous parents to CD child using the transmission disequilibrium test (TDT). Only alleles at two of the 57 markers had significant C^2 results ($p < 0.001$): IRF1p1 ($C^2=13.3$, $p=0.0003$) and D5S1984 ($C^2=14.0$, $p=0.0002$) (Table 1). A second stage of mapping was then undertaken to confirm, as well as to better delimit, the region of LD surrounding IRF1p1 and D5S1984. The development of new microsatellite markers was necessary. The marker with the most significant C^2 result was CAh17a ($C^2=16.2$, $p=0.00006$) and was located between IRF1p1 and D5S1984 (Table 2). Furthermore, the alleles 193, 156, 373, 140, 222, and 307 at markers GAh18a, IRF1p, CAh15a, CAh17a, D5S1984,

CSF2p10, respectively, define a haplotype conferring susceptibility to Crohn's disease (CD). In order to identify the sequence variant that would explain the genetic susceptibility to CD provided by this haplotype, a search was performed for all single nucleotide polymorphisms (SNPs) in this region of LD. The SNP discovery was
5 accomplished by direct sequencing of overlapping PCR products amplified from DNA samples from eight individuals (six CD patients, one unaffected family member, and one CEPH DNA as control). Table 3 shows the results of the SNP discovery analyses.

139 triads were genotyped for a total of 241 SNPs thus far, where at least 50 trios were fully genotyped. Using a C^2 value of 13 (corresponding to a p-value of 0.05) as
10 threshold, 12 SNPs were found to have a significant level of association with CD and extended over a region of 250 kb, from IRF1 to prolyl4 hydroxylase. These were markers IGR2055a_1, IGR2060a_1, IGR2063b_1, IGR2069a_2, IGR2078a_1, IGR2096a_1, IGR2198a_1, IGR2230a_1, IGR2277a_1, IGR3081a_1, IGR3096a_1, PROLYLex3_1 (see Table 4). Any of these best SNPs by themselves are in strong
15 association with CD and fully explain the microsatellite LD observations. Furthermore, the best SNPs have nearly identical association characteristics (that is, the allele at one SNP determines the allele of all others on any phased chromosome), confirming that a single risk haplotype extending approximately 250 kb is the source of all the observations of association in this region. Specifically, this haplotype is defined by the
20 alleles G, C, G, T, A, A, G, T, G, G, C, T at markers IGR2055a_1, IGR2060a_1, IGR2063b_1, IGR2069a_2, IGR2078a_1, IGR2096a_1, IGR2198a_1, IGR2230a_1, IGR2277a_1, IGR3081a_1, IGR3096a_1, PROLYLex3_1, respectively. The frequency of this haplotype is estimated to be approximately 37% in the general population. Furthermore, this haplotype is transmitted from heterozygous parents to CD patients at a
25 ratio of 2.5:1.

The invention relates to a isolated gene or nucleic acid molecule which comprises a single nucleotide polymorphism at a specific location. In a particular embodiment the invention relates to the variant allele of a gene having a single nucleotide polymorphism, which variant allele differs from a reference allele by one nucleotide at

the site(s) identified in Table 3. Complements of these nucleic acid segments are also included. The segments can be DNA or RNA, and can be double- or single-stranded. Segments can be, for example, 5-10, 5-15, 10-20, 5-25, 10-30, 10-50 or 10-100 bases long.

- 5 The invention further provides allele-specific oligonucleotides that hybridize to a gene comprising a single nucleotide polymorphism or to the complement of the gene. These oligonucleotides can be probes or primers.

10 The invention further provides a method of analyzing a nucleic acid from an individual. The method determines which base is present at any one of the polymorphic sites shown in Table 3. Optionally, a set of bases occupying a set of the polymorphic sites shown in Table 3 is determined. This type of analysis can be performed on a number of individuals, who are tested for the presence of a disease phenotype. The presence or absence of disease phenotype is then correlated with a base or set of bases present at the polymorphic site or sites in the individuals tested.

- 15 Thus, the invention further relates to a method of predicting the presence, absence, likelihood of the presence or absence, or severity of a particular phenotype or disorder associated with a particular genotype. The method comprises obtaining a nucleic acid sample from an individual and determining the identity of one or more bases (nucleotides) at polymorphic sites of genes described herein, wherein the presence of a particular base is correlated with a specified phenotype or disorder, thereby predicting
20 the presence, absence, likelihood of the presence or absence, or severity of the phenotype or disorder in the individual. In one embodiment of the invention, the phenotype is inflammatory bowel disease or Crohn's disease.

BRIEF DESCRIPTION OF THE DRAWING

- 25 The Figure shows multipoint nonparametric linkage results for the IBD genome scan. Multipoint LOD scores were calculated using the MAPMAKER/SIBS functions implemented in GENHUNTER 2.0. The thick black line indicates the LOD score along the length of each chromosome, and the tick marks indicate the position of the

microsatellite markers. The two horizontal lines depict the genome-wide thresholds for suggestive (LOD = 2.0) and significant linkage.

DETAILED DESCRIPTION OF THE INVENTION

Crohn's disease (CD) and ulcerative colitis (UC) are chronic, idiopathic
5 inflammatory disorders of the gastrointestinal tract. These inflammatory bowel diseases (IBD) have a peak incidence in early adulthood, and their combined prevalence is approximately 100-200/100,000. The inflammation in IBD is characterized by altered expression of both pro-inflammatory and immunoregulatory cytokines in the affected intestinal mucosa (Kmiec, *Arch Immunol Ther Expe (Warsz)* 46(3):143-155 (1998)).
10 Genetic factors are believed to play an important role, as the sibling risk (λ_s) calculated for IBD ranges from 15-40, with a stronger genetic contribution occurring for CD ($\lambda_s \sim 35$) as compared to UC ($\lambda_s \sim 15$). Additionally, relatives of individuals with IBD diagnosed at younger ages appear to be at an even higher risk.

CD is characterized by discontinuous, transmural inflammation affecting any part
15 of the gastrointestinal tract and is manifested by abdominal pain, chronic diarrhea, weight loss, bowel obstructions and fistulae. UC occurs as a continuous, mucosal inflammation affecting only the large intestine with primary symptoms including diarrhea, rectal bleeding and abdominal pain. The search for susceptibility genes for these two diseases has resulted in the identification of two potential susceptibility loci.
20 The first, called *IBD1*, is a CD-susceptibility locus that lies in the pericentromeric region of chromosome 16 (Hugo *et al.*, *Nature* 379:821-822 (1996)). The second (*IBD2*) is located in a 41 cM region surrounding marker D12S83 and appears to be linked to both CD and UC (Satsangi *et al.*, *Genetics* 14:199-202 (1996)). These putative loci, however, have only been replicated in some, but not all, subsequent
25 studies (Cavanaugh *et al.*, *Proc Natl Acad Sci USA* 62:291-298 (1998); Cho *et al.*, *The National Academy of Sciences* 95:7502-7507 (1998); Curren *et al.*, *Gastroenterology* 115:1-7 (1998); Duerr *et al.*, *The American Society of Human Genetics* 63:95-100 (1998); Rioux *et al.*, *Am. J. Hum. Genet.* 63:1086-1094 (1998); Yang *et al.*,

Gastroenterology 109:440-448 (1995)), supporting the belief that there exists substantial genetic heterogeneity. Furthermore, *IBD1* and *IBD2* only account for a fraction of the heritability of IBD, suggesting that additional loci contribute to disease susceptibility. Thus, as described herein, the susceptibility loci in a Canadian IBD
5 population was assessed by studying families with multiple affected siblings (McLeod *et al.*, *Dis Colon Rectum* 40:553-557 (1997)).

A genome-wide screen was performed on 181 IBD-affected sibling pairs (ASP) and 5 IBD-affected relative pairs (RP) from 163 families. Among these ASP, 122 were CD pairs, 25 were UC pairs, and 34 were "mixed" pairs (one sibling with either CD or
10 UC, the other with CD, UC or IC). All ASP and available parents (140 families had both parents available, 17 had one parent available, and 1 was missing both parents), as well as all RP, were genotyped with 312 microsatellite markers covering the genome with approximately 12 cM distance between markers. Simulations of this dataset indicated that the genome-wide threshold for suggestive linkage (the score expected to
15 occur one time at random in a genome scan) was at a LOD of 2.0. Using either this calculated threshold, or the published threshold of LOD 2.2 based on an infinitely dense map (Lander & Kruglyak, *Nature Genetics* 11:241-247 (1995)), multipoint nonparametric linkage analysis of these data revealed 4 loci which surpassed this threshold (Figure). Specifically, chromosome 3 had a peak LOD of 2.4 between
20 markers D3S1766 and D#S1285, chromosome 5 a peak LOD of 3.0 between GATA68A03 and D5S816, chromosome 6 a peak LOD of 2.3 between D6S1019 and D6S1017, and chromosome 19 a peak LOD of 4.6 between GATA21G05 and D19S586. In fact this chromosome 19 locus exceeds the threshold for genome-wide significance of 3.6 (Lander & Kruglyak, *Nature Genetics* 11:241-247 (1995)), and represents a novel
25 IBD susceptibility locus.

This novel locus maps to an extended region on 19p13 (Figure) that contains many different genes of immunologic interest such as intercellular adhesion molecule 1 (ICAM1), complement component 3 (C3), the thromboxane A2 receptor (TBXA2), leukotriene B4 hydroxylase (LTB4H), and the janus tyrosine kinases TYK2 and JAK3.

There is some evidence supporting their relevance in IBD susceptibility: 1) modest positive association results have been reported for the ICAM1 (Yang *et al.*, *Gastroenterology* 109:440-448 (1995)) and C3 molecules (Elmgreen *et al.*, *Acta Med Scand* 215(4):375-8 (1984)); 2) attempts to interfere with the TBXA2 (Taniguchi, 1997) and LTB4 (Hawkey *et al.*, *Agents Actions, Special Conference Issue* (1992)) mediated inflammatory pathways have shown some therapeutic value; and 3) the janus kinases have been shown to be important in the transduction of the molecular signal from cytokine receptors.

The finding of suggestive linkage to an approximately 30 cM region spanning the cytokine gene cluster on 5q31-q33, containing many of the immunoregulatory cytokines such as IL4, IL13, IL5 and IL3, led to the performance of higher density mapping in this region. Specifically, the original families and an additional 12 families were genotyped for 34 extra microsatellite markers. Multipoint nonparametric analysis were then performed using three different phenotypic categories: IBD, CD and CD16. In the first, all individuals with CD, UC or IC were designated as affected; in the second, only individuals with CD were designated as affected; in the third, only individuals with CD were designated as affected and only families with at least one affected sibling diagnosed at the age of 16 or younger were included. This last category was examined due to an expected enrichment for genetic factors over environmental causes. These analyses demonstrate the presence of a locus of genome-wide significance in the group with early onset CD (MLS = 3.9). Evidence for linkage to the syntenic region in mice has been reported in an induced model of colitis (Mahler, *Genomics* 55:147-156 (1999)).

Although the suggestive loci on chromosomes 3 and 6 identified as described herein have not yet been followed up with higher density mapping, it is important to note that the linkage peak on chromosome 3 is approximately 10 cM away from a previously reported suggestive locus (Satsangi *et al.*, *Nature Genetics* 14:199-202 (1996)), and the linkage peak on 6 lies approximately 20 cM centromeric to the major histocompatibility complex (MHC) Class II region. A recent study has described

linkage to this chromosome 6 region (Hampe *et al.*, *Am. J. Hum. Genet.* 64 (1999)), and a large meta-analysis of the results derived from 29 different studies has also reported that both CD and UC were associated with specific Class II alleles (Stokkers *et al.*, *Gut* 45:395-401 (1999)). Finally, in order to assess whether the *IBD1* and *IBD2* loci are
5 contributing to the IBD susceptibility in this population, exclusion mapping of the data was performed. These analyses demonstrate that the entire chromosome 12 can be excluded for loci of even modest effects ($\lambda_s > 1.5$), but can only loci conferring a $\lambda_s > 4$ on chromosome 16 can be excluded, suggesting that *IBD1* ($\lambda_s \sim 1.3$) could have gone undetected in the present study.

10 Thus, this work has identified two novel susceptibility loci: a locus on chromosome 5q31-33 that confers susceptibility to CD and a locus on chromosome 19p13 that confers susceptibility to IBD. Furthermore, particular SNPs within these loci have been identified which may be associated with disease susceptibility.

The present invention relates to a gene which comprises a single nucleotide
15 polymorphism (SNP) at a specific location. The gene which includes the SNP has at least two alleles, referred to herein as the reference allele and the variant allele. The reference allele (prototypical or wild type allele) has been designated arbitrarily and typically corresponds to the nucleotide sequence of the gene which has been deposited with GenBank or TIGR under a given Accession number. The variant allele differs
20 from the reference allele by one nucleotide at the site(s) identified in Table 3. The present invention also relates to variant alleles of the described genes and to complements of the variant alleles. The invention further relates to portions of the variant alleles and portions of complements of the variant alleles which comprise (encompass) the site of the SNP and are at least 5 nucleotides in length. Portions can
25 be, for example, 5-10, 5-15, 10-20, 5-25, 10-30, 10-50 or 10-100 bases long. For example, a portion of a variant allele which is 21 nucleotides in length includes the single nucleotide polymorphism (the nucleotide which differs from the reference allele at that site) and twenty additional nucleotides which flank the site in the variant allele. These nucleotides can be on one or both sides of the polymorphism.

Polymorphisms which are the subject of this invention are defined in Table 3. The reference sequence for many of the genes or gene fragments is provided in Table 5. For sequences which are not present in Table 5, the skilled artisan can readily determine the specific location of the polymorphism given the 3' and 5' nucleotide sequence
5 flanking the polymorphic site provided in Table 3 and the chromosomal loci information provided herein. The nucleotide sequences of the invention can be double- or single-stranded.

The invention further provides allele-specific oligonucleotides that hybridize to a gene comprising a single nucleotide polymorphism or to the complement of the gene.
10 These oligonucleotides can be probes or primers.

The invention further provides a method of analyzing a nucleic acid from an individual. The method determines which base is present at any one of the polymorphic sites shown in Table 3. Optionally, a set of bases occupying a set of the polymorphic sites shown in Table 3 is determined. This type of analysis can be performed on a
15 number of individuals, who are tested for the presence of a disease phenotype. The presence or absence of disease phenotype is then correlated with a base or set of bases present at the polymorphic site or sites in the individuals tested.

Thus, the invention further relates to a method of predicting the presence, absence, likelihood of the presence or absence, or severity of a particular phenotype or disorder
20 associated with a particular genotype. The method comprises obtaining a nucleic acid sample from an individual and determining the identity of one or more bases (nucleotides) at polymorphic sites of genes described herein, wherein the presence of a particular base is correlated with a specified phenotype or disorder, thereby predicting the presence, absence, likelihood of the presence or absence, or severity of the
25 phenotype or disorder in the individual.

DEFINITIONS

An oligonucleotide can be DNA or RNA, and single- or double-stranded. Oligonucleotides can be naturally occurring or synthetic, but are typically prepared by

synthetic means. Preferred oligonucleotides of the invention include segments of DNA, or their complements, which include any one of the polymorphic sites shown in Table 3. The segments can be between 5 and 250 bases, and, in specific embodiments, are between 5-10, 5-20, 10-20, 10-50, 20-50 or 10-100 bases. For example, the segment
5 can be 21 bases. The polymorphic site can occur within any position of the segment. The segments can be from any of the allelic forms of DNA shown in Table 3.

As used herein, the terms "nucleotide", "base" and "nucleic acid" are intended to be equivalent. The terms "nucleotide sequence", "nucleic acid sequence", "nucleic acid molecule" and "segment" are intended to be equivalent.

10 Hybridization probes are oligonucleotides which bind in a base-specific manner to a complementary strand of nucleic acid. Such probes include peptide nucleic acids, as described in Nielsen *et al.*, *Science* 254, 1497-1500 (1991). Probes can be any length suitable for specific hybridization to the target nucleic acid sequence. The most appropriate length of the probe may vary depending upon the hybridization method in
15 which it is being used; for example, particular lengths may be more appropriate for use in microfabricated arrays, while other lengths may be more suitable for use in classical hybridization methods. Such optimizations are known to the skilled artisan. Suitable probes and primers can range from about 5 nucleotides to about 30 nucleotides in length. For example, probes and primers can be 5, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24,
20 25, 26, 28 or 30 nucleotides in length. The probe or primer preferably overlaps at least one polymorphic site occupied by any of the possible variant nucleotides. The nucleotide sequence can correspond to the coding sequence of the allele or to the complement of the coding sequence of the allele.

As used herein, the term "primer" refers to a single-stranded oligonucleotide
25 which acts as a point of initiation of template-directed DNA synthesis under appropriate conditions (*e.g.*, in the presence of four different nucleoside triphosphates and an agent for polymerization, such as DNA or RNA polymerase or reverse transcriptase) in an appropriate buffer and at a suitable temperature. The appropriate length of a primer depends on the intended use of the primer, but typically ranges from 15 to 30

nucleotides. Short primer molecules generally require cooler temperatures to form sufficiently stable hybrid complexes with the template. A primer need not reflect the exact sequence of the template, but must be sufficiently complementary to hybridize with a template. The term primer site refers to the area of the target DNA to which a
5 primer hybridizes. The term primer pair refers to a set of primers including a 5' (upstream) primer that hybridizes with the 5' end of the DNA sequence to be amplified and a 3' (downstream) primer that hybridizes with the complement of the 3' end of the sequence to be amplified.

As used herein, linkage describes the tendency of genes, alleles, loci or genetic
10 markers to be inherited together as a result of their location on the same chromosome. It can be measured by percent recombination between the two genes, alleles, loci or genetic markers.

As used herein, polymorphism refers to the occurrence of two or more genetically determined alternative sequences or alleles in a population. A polymorphic marker or
15 site is the locus at which divergence occurs. Preferred markers have at least two alleles, each occurring at frequency of greater than 1%, and more preferably greater than 10% or 20% of a selected population. A polymorphic locus may be as small as one base pair. Polymorphic markers include restriction fragment length polymorphisms, variable number of tandem repeats (VNTR's), hypervariable regions, minisatellites, dinucleotide
20 repeats, trinucleotide repeats, tetranucleotide repeats, simple sequence repeats, and insertion elements such as Alu. The first identified allelic form is arbitrarily designated as the reference form and other allelic forms are designated as alternative or variant alleles. The allelic form occurring most frequently in a selected population is sometimes referred to as the wildtype form. Diploid organisms may be homozygous or
25 heterozygous for allelic forms. A diallelic or biallelic polymorphism has two forms. A triallelic polymorphism has three forms.

Work described herein pertains to the resequencing of large numbers of genes in a large number of individuals to identify polymorphisms which can predispose individuals to disease, particularly IBD.

By altering amino acid sequence, SNPs may alter the function of the encoded proteins. The discovery of the SNP facilitates biochemical analysis of the variants and the development of assays to characterize the variants and to screen for pharmaceutical that would interact directly with on or another form of the protein. SNPs (including
5 silent SNPs) may also alter the regulation of the gene at the transcriptional or post-transcriptional level. SNPs (including silent SNPs) also enable the development of specific DNA, RNA, or protein-based diagnostics that detect the presence or absence of the polymorphism in particular conditions.

A single nucleotide polymorphism occurs at a polymorphic site occupied by a
10 single nucleotide, which is the site of variation between allelic sequences. The site is usually preceded by and followed by highly conserved sequences of the allele (e.g., sequences that vary in less than 1/100 or 1/1000 members of the populations).

A single nucleotide polymorphism usually arises due to substitution of one nucleotide for another at the polymorphic site. A transition is the replacement of one
15 purine by another purine or one pyrimidine by another pyrimidine. A transversion is the replacement of a purine by a pyrimidine or vice versa. Single nucleotide polymorphisms can also arise from a deletion of a nucleotide or an insertion of a nucleotide relative to a reference allele. Typically the polymorphic site is occupied by a base other than the reference base. For example, where the reference allele contains the
20 base "T" at the polymorphic site, the altered allele can contain a "C", "G" or "A" at the polymorphic site.

Hybridizations are usually performed under stringent conditions, for example, at a salt concentration of no more than 1 M and a temperature of at least 25°C. For example, conditions of 5X SSPE (750 mM NaCl, 50 mM NaPhosphate, 5 mM EDTA,
25 pH 7.4) and a temperature of 25-30°C, or equivalent conditions, are suitable for allele-specific probe hybridizations. Equivalent conditions can be determined by varying one or more of the parameters given as an example, as known in the art, while maintaining a similar degree of identity or similarity between the target nucleotide sequence and the primer or probe used.

The term "isolated" is used herein to indicate that the material in question exists in a physical milieu distinct from that in which it occurs in nature. For example, an isolated nucleic acid of the invention may be substantially isolated with respect to the complex cellular milieu in which it naturally occurs. In some instances, the isolated material will form part of a composition (for example, a crude extract containing other substances), buffer system or reagent mix. In other circumstance, the material may be purified to essential homogeneity, for example as determined by PAGE or column chromatography such as HPLC. Preferably, an isolated nucleic acid comprises at least about 50, 80 or 90 percent (on a molar basis) of all macromolecular species present.

10 I. Novel Polymorphisms of the Invention

The novel polymorphisms of the invention are shown in Table 3.

II. Analysis of Polymorphisms

A. Preparation of Samples

Polymorphisms are detected in a target nucleic acid from an individual being analyzed. For assay of genomic DNA, virtually any biological sample (other than pure red blood cells) is suitable. For example, convenient tissue samples include whole blood, semen, saliva, tears, urine, fecal material, sweat, buccal, skin and hair. For assay of cDNA or mRNA, the tissue sample must be obtained from an organ in which the target nucleic acid is expressed. For example, if the target nucleic acid is a cytochrome P450, the liver is a suitable source.

Many of the methods described below require amplification of DNA from target samples. This can be accomplished by e.g., PCR. *See generally PCR Technology: Principles and Applications for DNA Amplification* (ed. H.A. Erlich, Freeman Press, NY, NY, 1992); *PCR Protocols: A Guide to Methods and Applications* (eds. Innis, et al., Academic Press, San Diego, CA, 1990); Mattila et al., *Nucleic Acids Res.* 19, 4967 (1991); Eckert et al., *PCR Methods and Applications* 1, 17 (1991); PCR (eds. McPherson et al., IRL Press, Oxford); and U.S. Patent 4,683,202.

Other suitable amplification methods include the ligase chain reaction (LCR) (see Wu and Wallace, *Genomics* 4, 560 (1989), Landegren *et al.*, *Science* 241, 1077 (1988), transcription amplification (Kwoh *et al.*, *Proc. Natl. Acad. Sci. USA* 86, 1173 (1989)), and self-sustained sequence replication (Guatelli *et al.*, *Proc. Nat. Acad. Sci. USA*, 87, 1874 (1990)) and nucleic acid based sequence amplification (NASBA). The latter two
5 amplification methods involve isothermal reactions based on isothermal transcription, which produce both single stranded RNA (ssRNA) and double stranded DNA (dsDNA) as the amplification products in a ratio of about 30 or 100 to 1, respectively.

B. Detection of Polymorphisms in Target DNA

10 The polymorphisms identified as described herein can be used as a platform for genotyping (i.e., determining the genotype of) individuals. This analysis determines which form(s) of a characterized (known) polymorphism are present in individuals under test. There are a variety of suitable procedures, which are discussed in turn.

1. Allele-Specific Probes

15 The design and use of allele-specific probes for analyzing polymorphisms is described by e.g., Saiki *et al.*, *Nature* 324, 163-166 (1986); Dattagupta, EP 235,726, Saiki, WO 89/11548. Allele-specific probes can be designed that hybridize to a segment of target DNA from one individual but do not hybridize to the corresponding segment from another individual due to the presence of different polymorphic forms in
20 the respective segments from the two individuals. Hybridization conditions should be sufficiently stringent that there is a significant difference in hybridization intensity between alleles, and preferably an essentially binary response, whereby a probe hybridizes to only one of the alleles. Some probes are designed to hybridize to a segment of target DNA such that the polymorphic site aligns with a central position
25 (e.g., in a 15-mer at the 7 position; in a 16-mer, at either the 8 or 9 position) of the probe. This design of probe achieves good discrimination in hybridization between different allelic forms.

Allele-specific probes are often used in pairs, one member of a pair showing a perfect match to a reference form of a target sequence and the other member showing a perfect match to a variant form. Several pairs of probes can then be immobilized on the same support for simultaneous analysis of multiple polymorphisms within the same
5 target sequence.

2. Tiling Arrays

The polymorphisms can also be identified by hybridization to nucleic acid arrays, some examples of which are described in WO 95/11995. One form of such arrays is described in the Examples section in connection with de novo identification of
10 polymorphisms. The same array or a different array can be used for analysis of characterized polymorphisms. WO 95/11995 also describes subarrays that are optimized for detection of a variant form of a precharacterized polymorphism. Such a subarray contains probes designed to be complementary to a second reference sequence, which is an allelic variant of the first reference sequence. The second group of probes is
15 designed by the same principles as described in the Examples, except that the probes exhibit complementarity to the second reference sequence. The inclusion of a second group (or further groups) can be particularly useful for analyzing short subsequences of the primary reference sequence in which multiple mutations are expected to occur within a short distance commensurate with the length of the probes (e.g., two or more
20 mutations within 9 to 21 bases).

3. Allele-Specific Primers

An allele-specific primer hybridizes to a site on target DNA overlapping a polymorphism and only primes amplification of an allelic form to which the primer exhibits perfect complementarity. See Gibbs, *Nucleic Acid Res.* 17, 2427-2448 (1989).
25 This primer is used in conjunction with a second primer which hybridizes at a distal site. Amplification proceeds from the two primers, resulting in a detectable product which indicates the particular allelic form is present. A control is usually performed with a

second pair of primers, one of which shows a single base mismatch at the polymorphic site and the other of which exhibits perfect complementarity to a distal site. The single-base mismatch prevents amplification and no detectable product is formed. The method works best when the mismatch is included in the 3'-most position of the oligonucleotide
5 aligned with the polymorphism because this position is most destabilizing to elongation from the primer (see, e.g., WO 93/22456).

4. Direct-Sequencing

The direct analysis of the sequence of polymorphisms of the present invention can be accomplished using either the dideoxy chain termination method or the Maxam -
10 Gilbert method (see Sambrook *et al.*, *Molecular Cloning, A Laboratory Manual* (2nd Ed., CSHP, New York 1989); Zyskind *et al.*, *Recombinant DNA Laboratory Manual*, (Acad. Press, 1988)).

5. Denaturing Gradient Gel Electrophoresis

Amplification products generated using the polymerase chain reaction can be
15 analyzed by the use of denaturing gradient gel electrophoresis. Different alleles can be identified based on the different sequence-dependent melting properties and electrophoretic migration of DNA in solution. Erlich, ed., *PCR Technology, Principles and Applications for DNA Amplification*, (W.H. Freeman and Co, New York, 1992), Chapter 7.

20 6. Single-Strand Conformation Polymorphism Analysis

Alleles of target sequences can be differentiated using single-strand conformation polymorphism analysis, which identifies base differences by alteration in electrophoretic migration of single stranded PCR products, as described in Orita *et al.*, *Proc. Nat. Acad. Sci.* 86, 2766-2770 (1989). Amplified PCR products can be generated as described
25 above, and heated or otherwise denatured, to form single stranded amplification products. Single-stranded nucleic acids may refold or form secondary structures which

are partially dependent on the base sequence. The different electrophoretic mobilities of single-stranded amplification products can be related to base-sequence differences between alleles of target sequences.

7. Single Base Extension

- 5 An alternative method for identifying and analyzing polymorphisms is based on single-base extension (SBE) of a fluorescently-labeled primer coupled with fluorescence resonance energy transfer (FRET) between the label of the added base and the label of the primer. Typically, the method, such as that described by Chen *et al.*, (*PNAS* 94:10756-61 (1997), incorporated herein by reference) uses a locus-specific
- 10 oligonucleotide primer labeled on the 5' terminus with 5-carboxyfluorescein (FAM). This labeled primer is designed so that the 3' end is immediately adjacent to the polymorphic site of interest. The labeled primer is hybridized to the locus, and single base extension of the labeled primer is performed with fluorescently labeled dideoxyribonucleotides (ddNTPs) in dye-terminator sequencing fashion, except that no
- 15 deoxyribonucleotides are present. An increase in fluorescence of the added ddNTP in response to excitation at the wavelength of the labeled primer is used to infer the identity of the added nucleotide.

III. Methods of Use

- After determining polymorphic form(s) present in an individual at one or more
- 20 polymorphic sites, this information can be used in a number of methods.

A. Forensics

- Determination of which polymorphic forms occupy a set of polymorphic sites in an individual identifies a set of polymorphic forms that distinguishes the individual. *See generally* National Research Council, *The Evaluation of Forensic DNA Evidence* (Eds.
- 25 Pollard *et al.*, National Academy Press, DC, 1996). The more sites that are analyzed, the lower the probability that the set of polymorphic forms in one individual is the same

as that in an unrelated individual. Preferably, if multiple sites are analyzed, the sites are unlinked. Thus, polymorphisms of the invention are often used in conjunction with polymorphisms in distal genes. Preferred polymorphisms for use in forensics are biallelic because the population frequencies of two polymorphic forms can usually be
5 determined with greater accuracy than those of multiple polymorphic forms at multi-allelic loci.

The capacity to identify a distinguishing or unique set of forensic markers in an individual is useful for forensic analysis. For example, one can determine whether a blood sample from a suspect matches a blood or other tissue sample from a crime scene
10 by determining whether the set of polymorphic forms occupying selected polymorphic sites is the same in the suspect and the sample. If the set of polymorphic markers does not match between a suspect and a sample, it can be concluded (barring experimental error) that the suspect was not the source of the sample. If the set of markers does match, one can conclude that the DNA from the suspect is consistent with that found at
15 the crime scene. If frequencies of the polymorphic forms at the loci tested have been determined (e.g., by analysis of a suitable population of individuals), one can perform a statistical analysis to determine the probability that a match of suspect and crime scene sample would occur by chance.

$p(ID)$ is the probability that two random individuals have the same polymorphic or
20 allelic form at a given polymorphic site. In biallelic loci, four genotypes are possible: AA, AB, BA, and BB. If alleles A and B occur in a haploid genome of the organism with frequencies x and y , the probability of each genotype in a diploid organism is (see WO 95/12607):

Homozygote: $p(AA) = x^2$
25 Homozygote: $p(BB) = y^2 = (1-x)^2$
Single Heterozygote: $p(AB) = p(BA) = xy = x(1-x)$
Both Heterozygotes: $p(AB+BA) = 2xy = 2x(1-x)$

The probability of identity at one locus (i.e, the probability that two individuals, picked at random from a population will have identical polymorphic forms at a given locus) is given by the equation:

$$p(\text{ID}) = (x^2)^2 + (2xy)^2 + (y^2)^2.$$

- 5 These calculations can be extended for any number of polymorphic forms at a given locus. For example, the probability of identity $p(\text{ID})$ for a 3-allele system where the alleles have the frequencies in the population of x , y and z , respectively, is equal to the sum of the squares of the genotype frequencies:

$$p(\text{ID}) = x^4 + (2xy)^2 + (2yz)^2 + (2xz)^2 + z^4 + y^4$$

- 10 In a locus of n alleles, the appropriate binomial expansion is used to calculate $p(\text{ID})$ and $p(\text{exc})$.

The cumulative probability of identity ($\text{cum } p(\text{ID})$) for each of multiple unlinked loci is determined by multiplying the probabilities provided by each locus.

$$\text{cum } p(\text{ID}) = p(\text{ID}1)p(\text{ID}2)p(\text{ID}3).... p(\text{ID}n)$$

- 15 The cumulative probability of non-identity for n loci (i.e. the probability that two random individuals will be different at 1 or more loci) is given by the equation:

$$\text{cum } p(\text{nonID}) = 1 - \text{cum } p(\text{ID}).$$

- If several polymorphic loci are tested, the cumulative probability of non-identity for random individuals becomes very high (e.g., one billion to one). Such probabilities
20 can be taken into account together with other evidence in determining the guilt or innocence of the suspect.

B. Paternity Testing

- The object of paternity testing is usually to determine whether a male is the father of a child. In most cases, the mother of the child is known and thus, the mother's
25 contribution to the child's genotype can be traced. Paternity testing investigates whether the part of the child's genotype not attributable to the mother is consistent with that of

the putative father. Paternity testing can be performed by analyzing sets of polymorphisms in the putative father and the child.

If the set of polymorphisms in the child attributable to the father does not match the set of polymorphisms of the putative father, it can be concluded, barring
 5 experimental error, that the putative father is not the real father. If the set of polymorphisms in the child attributable to the father does match the set of polymorphisms of the putative father, a statistical calculation can be performed to determine the probability of coincidental match.

The probability of parentage exclusion (representing the probability that a random
 10 male will have a polymorphic form at a given polymorphic site that makes him incompatible as the father) is given by the equation (see WO 95/12607):

$$p(\text{exc}) = xy(1-xy)$$

where x and y are the population frequencies of alleles A and B of a biallelic polymorphic site.

15 (At a triallelic site $p(\text{exc}) = xy(1-xy) + yz(1-yz) + xz(1-xz) + 3xyz(1-xyz)$), where x, y and z are the respective population frequencies of alleles A, B and C).

The probability of non-exclusion is

$$p(\text{non-exc}) = 1 - p(\text{exc})$$

The cumulative probability of non-exclusion (representing the value obtained
 20 when n loci are used) is thus:

$$\text{cum } p(\text{non-exc}) = p(\text{non-exc1})p(\text{non-exc2})p(\text{non-exc3})\dots p(\text{non-excn})$$

The cumulative probability of exclusion for n loci (representing the probability that a random male will be excluded)

$$\text{cum } p(\text{exc}) = 1 - \text{cum } p(\text{non-exc}).$$

25 If several polymorphic loci are included in the analysis, the cumulative probability of exclusion of a random male is very high. This probability can be taken into account in assessing the liability of a putative father whose polymorphic marker set matches the child's polymorphic marker set attributable to his/her father.

C. Correlation of Polymorphisms with Phenotypic Traits

The polymorphisms of the invention may contribute to the phenotype of an organism in different ways. Some polymorphisms occur within a protein coding sequence and contribute to phenotype by affecting protein structure. The effect may be neutral, beneficial or detrimental, or both beneficial and detrimental, depending on the circumstances. For example, a heterozygous sickle cell mutation confers resistance to malaria, but a homozygous sickle cell mutation is usually lethal. Other polymorphisms occur in noncoding regions but may exert phenotypic effects indirectly via influence on replication, transcription, and translation. A single polymorphism may affect more than one phenotypic trait. Likewise, a single phenotypic trait may be affected by polymorphisms in different genes. Further, some polymorphisms predispose an individual to a distinct mutation that is causally related to a certain phenotype. For example, the polymorphisms identified herein and shown in Table 3 are present in the chromosomal loci which have been identified as described herein as conferring susceptibility to IBD such as CD and UC.

Correlation is performed for a population of individuals who have been tested for the presence or absence of a phenotypic trait of interest and for polymorphic markers sets. To perform such analysis, the presence or absence of a set of polymorphisms (i.e. a polymorphic set) is determined for a set of the individuals, some of whom exhibit a particular trait, and some of which exhibit lack of the trait. The alleles of each polymorphism of the set are then reviewed to determine whether the presence or absence of a particular allele is associated with the trait of interest. Correlation can be performed by standard statistical methods such as a χ -squared test and statistically significant correlations between polymorphic form(s) and phenotypic characteristics are noted. For example, it might be found that the presence of allele A1 at polymorphism A correlates with heart disease. As a further example, it might be found that the combined presence of allele A1 at polymorphism A and allele B1 at polymorphism B correlates with increased susceptibility to IBD (e.g., CD and UC).

Such correlations can be exploited in several ways. In the case of a strong correlation between a set of one or more polymorphic forms and a disease for which treatment is available, detection of the polymorphic form set in a human or animal patient may justify immediate administration of treatment, or at least the institution of regular monitoring of the patient. Detection of a polymorphic form correlated with serious disease in a couple contemplating a family may also be valuable to the couple in their reproductive decisions. For example, the female partner might elect to undergo *in vitro* fertilization to avoid the possibility of transmitting such a polymorphism from her husband to her offspring. In the case of a weaker, but still statistically significant correlation between a polymorphic set and human disease, immediate therapeutic intervention or monitoring may not be justified. Nevertheless, the patient can be motivated to begin simple life-style changes (e.g., diet, exercise) that can be accomplished at little cost to the patient but confer potential benefits in reducing the risk of conditions to which the patient may have increased susceptibility by virtue of variant alleles. Identification of a polymorphic set in a patient correlated with enhanced receptiveness to one of several treatment regimes for a disease indicates that this treatment regime should be followed.

For animals and plants, correlations between characteristics and phenotype are useful for breeding for desired characteristics. For example, Beitz *et al.*, US 5,292,639 discuss use of bovine mitochondrial polymorphisms in a breeding program to improve milk production in cows. To evaluate the effect of mtDNA D-loop sequence polymorphism on milk production, each cow was assigned a value of 1 if variant or 0 if wildtype with respect to a prototypical mitochondrial DNA sequence at each of 17 locations considered. Each production trait was analyzed individually with the following animal model:

$$Y_{ijkpn} = \mu + YS_i + P_j + X_k + \beta_1 + \dots \beta_{17} + PE_n + a_n + e_p$$

where Y_{ijkpn} is the milk, fat, fat percentage, SNF, SNF percentage, energy concentration, or lactation energy record; μ is an overall mean; YS_i is the effect common to all cows calving in year-season; X_k is the effect common to cows in either the high or average

selection line; β_1 to β_{17} are the binomial regressions of production record on mtDNA D-loop sequence polymorphisms; PE_n is permanent environmental effect common to all records of cow n ; a_n is effect of animal n and is composed of the additive genetic contribution of sire and dam breeding values and a Mendelian sampling effect; and e_p is a random residual. It was found that eleven of seventeen polymorphisms tested influenced at least one production trait. Bovines having the best polymorphic forms for milk production at these eleven loci are used as parents for breeding the next generation of the herd.

D. Genetic Mapping of Phenotypic Traits

The previous section concerns identifying correlations between phenotypic traits (e.g., IBD) and polymorphisms that directly or indirectly contribute to those traits, such as those identified in Table 3. The present section describes identification of a physical linkage between a genetic locus associated with a trait of interest and polymorphic markers that are not associated with the trait, but are in physical proximity with the genetic locus responsible for the trait and co-segregate with it. Such analysis is useful for mapping a genetic locus associated with a phenotypic trait to a chromosomal position, and thereby cloning gene(s) responsible for the trait. See Lander *et al.*, *Proc. Natl. Acad. Sci. (USA)* 83, 7353-7357 (1986); Lander *et al.*, *Proc. Natl. Acad. Sci. (USA)* 84, 2363-2367 (1987); Donis-Keller *et al.*, *Cell* 51, 319-337 (1987); Lander *et al.*, *Genetics* 121, 185-199 (1989)). Genes localized by linkage can be cloned by a process known as directional cloning. See Wainwright, *Med. J. Australia* 159, 170-174 (1993); Collins, *Nature Genetics* 1, 3-6 (1992).

Linkage studies are typically performed on members of a family. Available members of the family are characterized for the presence or absence of a phenotypic trait and for a set of polymorphic markers. The distribution of polymorphic markers in an informative meiosis is then analyzed to determine which polymorphic markers co-segregate with a phenotypic trait. See, e.g., Kerem *et al.*, *Science* 245, 1073-1080

(1989); Monaco *et al.*, *Nature* 316, 842 (1985); Yamoka *et al.*, *Neurology* 40, 222-226 (1990); Rossiter *et al.*, *FASEB Journal* 5, 21-27 (1991).

Linkage is analyzed by calculation of LOD (log of the odds) values. A lod value is the relative likelihood of obtaining observed segregation data for a marker and a genetic locus when the two are located at a recombination fraction θ , versus the situation in which the two are not linked, and thus segregating independently (Thompson & Thompson, *Genetics in Medicine* (5th ed, W.B. Saunders Company, Philadelphia, 1991); Strachan, "Mapping the human genome" in *The Human Genome* (BIOS Scientific Publishers Ltd, Oxford), Chapter 4). A series of likelihood ratios are calculated at various recombination fractions (θ), ranging from $\theta = 0.0$ (coincident loci) to $\theta = 0.50$ (unlinked). Thus, the likelihood at a given value of θ is: probability of data if loci linked at θ to probability of data if loci unlinked. The computed likelihoods are usually expressed as the \log_{10} of this ratio (i.e., a lod score). For example, a lod score of 3 indicates 1000:1 odds against an apparent observed linkage being a coincidence.

The use of logarithms allows data collected from different families to be combined by simple addition. Computer programs are available for the calculation of lod scores for differing values of θ (e.g., LIPED, MLINK (Lathrop, *Proc. Nat. Acad. Sci. (USA)* 81, 3443-3446 (1984)). For any particular lod score, a recombination fraction may be determined from mathematical tables. See Smith *et al.*, *Mathematical tables for research workers in human genetics* (Churchill, London, 1961); Smith, *Ann. Hum. Genet.* 32, 127-150 (1968). The value of θ at which the lod score is the highest is considered to be the best estimate of the recombination fraction.

Positive lod score values suggest that the two loci are linked, whereas negative values suggest that linkage is less likely (at that value of θ) than the possibility that the two loci are unlinked. By convention, a combined lod score of +3 or greater (equivalent to greater than 1000:1 odds in favor of linkage) is considered definitive evidence that two loci are linked. Similarly, by convention, a negative lod score of -2 or less is taken as definitive evidence against linkage of the two loci being compared. Negative linkage

data are useful in excluding a chromosome or a segment thereof from consideration. The search focuses on the remaining non-excluded chromosomal locations.

IV. Modified Polypeptides and Gene Sequences

The invention further provides variant forms of nucleic acids and corresponding
5 proteins. The nucleic acids comprise one of the sequences described in Table 3, in which the polymorphic position is occupied by one of the alternative bases for that position. Some nucleic acids encode full-length variant forms of proteins. Similarly, variant proteins have the prototypical amino acid sequences encoded by nucleic acid sequences shown in Table 3, (read so as to be in-frame with the full-length coding
10 sequence of which it is a component) except at an amino acid encoded by a codon including one of the polymorphic positions shown in Table 3. That position is occupied by the amino acid coded by the corresponding codon in any of the alternative forms shown in Table 3.

Variant genes can be expressed in an expression vector in which a variant gene is
15 operably linked to a native or other promoter. Usually, the promoter is a eukaryotic promoter for expression in a mammalian cell. The transcription regulation sequences typically include a heterologous promoter and optionally an enhancer which is recognized by the host. The selection of an appropriate promoter, for example trp, lac, phage promoters, glycolytic enzyme promoters and tRNA promoters, depends on the
20 host selected. Commercially available expression vectors can be used. Vectors can include host-recognized replication systems, amplifiable genes, selectable markers, host sequences useful for insertion into the host genome, and the like.

The means of introducing the expression construct into a host cell varies depending upon the particular construction and the target host. Suitable means include
25 fusion, conjugation, transfection, transduction, electroporation or injection, as described in Sambrook, *supra*. A wide variety of host cells can be employed for expression of the variant gene, both prokaryotic and eukaryotic. Suitable host cells include bacteria such as *E. coli*, yeast, filamentous fungi, insect cells, mammalian cells, typically

immortalized, *e.g.*, mouse, CHO, human and monkey cell lines and derivatives thereof. Preferred host cells are able to process the variant gene product to produce an appropriate mature polypeptide. Processing includes glycosylation, ubiquitination, disulfide bond formation, general post-translational modification, and the like. As used
5 herein, "gene product" includes mRNA, peptide and protein products.

The protein may be isolated by conventional means of protein biochemistry and purification to obtain a substantially pure product, *i.e.*, 80, 95 or 99% free of cell component contaminants, as described in Jacoby, *Methods in Enzymology* Volume 104, Academic Press, New York (1984); Scopes, *Protein Purification, Principles and*
10 *Practice*, 2nd Edition, Springer-Verlag, New York (1987); and Deutscher (ed), *Guide to Protein Purification, Methods in Enzymology*, Vol. 182 (1990). If the protein is secreted, it can be isolated from the supernatant in which the host cell is grown. If not secreted, the protein can be isolated from a lysate of the host cells.

The invention further provides transgenic nonhuman animals capable of
15 expressing an exogenous variant gene and/or having one or both alleles of an endogenous variant gene inactivated. Expression of an exogenous variant gene is usually achieved by operably linking the gene to a promoter and optionally an enhancer, and microinjecting the construct into a zygote. See Hogan *et al.*, "Manipulating the Mouse Embryo, A Laboratory Manual," Cold Spring Harbor Laboratory. Inactivation
20 of endogenous variant genes can be achieved by forming a transgene in which a cloned variant gene is inactivated by insertion of a positive selection marker. See Capecchi, *Science* 244, 1288-1292 (1989). The transgene is then introduced into an embryonic stem cell, where it undergoes homologous recombination with an endogenous variant gene. Mice and other rodents are preferred animals. Such animals provide useful drug
25 screening systems.

In addition to substantially full-length polypeptides expressed by variant genes, the present invention includes biologically active fragments of the polypeptides, or analogs thereof, including organic molecules which simulate the interactions of the peptides. Biologically active fragments include any portion of the full-length

polypeptide which confers a biological function on the variant gene product, including ligand binding, and antibody binding. Ligand binding includes binding by nucleic acids, proteins or polypeptides, small biologically active molecules, or large cellular structures.

Polyclonal and/or monoclonal antibodies that specifically bind to variant gene products but not to corresponding prototypical gene products are also provided.

Antibodies can be made by injecting mice or other animals with the variant gene product or synthetic peptide fragments thereof. Monoclonal antibodies are screened as are described, for example, in Harlow & Lane, *Antibodies, A Laboratory Manual*, Cold Spring Harbor Press, New York (1988); Goding, *Monoclonal antibodies, Principles and Practice* (2d ed.) Academic Press, New York (1986). Monoclonal antibodies are tested for specific immunoreactivity with a variant gene product and lack of immunoreactivity to the corresponding prototypical gene product. These antibodies are useful in diagnostic assays for detection of the variant form, or as an active ingredient in a pharmaceutical composition.

15 V. Kits

The invention further provides kits comprising at least one allele-specific oligonucleotide as described herein. Often, the kits contain one or more pairs of allele-specific oligonucleotides hybridizing to different forms of a polymorphism. In some kits, the allele-specific oligonucleotides are provided immobilized to a substrate. For example, the same substrate can comprise allele-specific oligonucleotide probes for detecting at least 10, 100 or all of the polymorphisms shown in Table 3. Optional additional components of the kit include, for example, restriction enzymes, reverse-transcriptase or polymerase, the substrate nucleoside triphosphates, means used to label (for example, an avidin-enzyme conjugate and enzyme substrate and chromogen if the label is biotin), and the appropriate buffers for reverse transcription, PCR, or hybridization reactions. Usually, the kit also contains instructions for carrying out the methods.

The following Examples are offered for the purpose of illustrating the present invention and are not to be construed to limit the scope of this invention. The teachings of all references cited herein are hereby incorporated herein by reference.

EXAMPLES

5 With the goal of identifying IBD susceptibility genes, a genomewide scan was undertaken in 163 pedigrees, and three regions of suggestive linkage (3, 5q31-33, 6p) and one of significant linkage to 19p13 (LOD = 4.6) were identified. Higher density mapping in the suggestive 5q31-33 region revealed a CD susceptibility locus of genome-wide significance (LOD = 3.9). Importantly, the 5q31-p33 localizes to the
10 major immunoregulatory cytokine gene cluster and the 19p13 locus to a region containing numerous genes encoding cytokine/chemokine receptors and other inflammatory-associated molecules that could have a direct role in disease susceptibility.

 In order to pursue the evidence of linkage to chromosome 5, a systematic linkage
15 disequilibrium (LD) approach was adopted. The approach that was used in the first stage of LD mapping was to genotype all known microsatellite markers in the 18 cM between D5S1435 and D5S1480, as these two markers delimit a region of a 2 LOD drop on either side of the linkage peak centered at marker D5S2497. All alleles for each marker were examined for evidence of excess transmission from heterozygous parents
20 to CD child using the transmission disequilibrium test (TDT). Only alleles at two of the 57 markers had significant C^2 results ($p < 0.001$): IRF1p1 ($C^2 = 13.3$, $p = 0.0003$) and D5S1984 ($C^2 = 14.0$, $p = 0.0002$) (Table 1). A second stage of mapping was then undertaken to confirm, as well as to better delimit, the region of LD surrounding IRF1p1 and D5S1984. The development of new microsatellite markers was necessary.
25 The marker with the most significant C^2 result was CAh17a ($C^2 = 16.2$, $p = 0.00006$) and was located between IRF1p1 and D5S1984 (Table 2). Furthermore, the alleles 193, 156, 373, 140, 222, and 307 at markers GAh18a, IRF1p, CAh15a, CAh17a, D5S1984, CSF2p10, respectively, define a haplotype conferring susceptibility to Crohn's disease

(CD). In order to identify the sequence variant that would explain the genetic susceptibility to CD provided by this haplotype, a search was performed for all single nucleotide polymorphisms (SNPs) in this region of LD. The SNP discovery was accomplished by direct sequencing of overlapping PCR products amplified from DNA samples from eight individuals (six CD patients, one unaffected family member, and one CEPH DNA as control). Table 3 shows the results of the SNP discovery analyses.

139 triads were genotyped for a total of 241 SNPs thus far, where at least 50 trios were fully genotyped. Using a C^2 value of 13 (corresponding to a p-value of 0.05) as threshold, 12 SNPs were found to have a significant level of association with CD and extended over a region of 250 kb, from IRF1 to prolyl4 hydroxylase. These were markers IGR2055a_1, IGR2060a_1, IGR2063b_1, IGR2069a_2, IGR2078a_1, IGR2096a_1, IGR2198a_1, IGR2230a_1, IGR2277a_1, IGR3081a_1, IGR3096a_1, PROLYLex3_1 (see Table 4). Any of these best SNPs by themselves are in strong association with CD and fully explain the microsatellite LD observations. Furthermore, the best SNPs have nearly identical association characteristics (that is, the allele at one SNP determines the allele of all others on any phased chromosome), confirming that a single risk haplotype extending approximately 250 kb is the source of all the observations of association in this region. Specifically, this haplotype is defined by the alleles G, C, G, T, A, A, G, T, G, G, C, T at markers IGR2055a_1, IGR2060a_1, IGR2063b_1, IGR2069a_2, IGR2078a_1, IGR2096a_1, IGR2198a_1, IGR2230a_1, IGR2277a_1, IGR3081a_1, IGR3096a_1, PROLYLex3_1, respectively. The frequency of this haplotype is estimated to be approximately 37% in the general population. Furthermore, this haplotype is transmitted from heterozygous parents to CD patients at a ratio of 2.5:1.

25 Families

For the linkage study, multicase families with 2 or more siblings affected by IBD were identified by review of clinical charts of all patients registered in the Mount Sinai

Hospital Inflammatory Bowel Disease Unit patient database and from the Hospital for Sick Children IBD database. Patients were also referred by physicians in the Greater Toronto Area (GTA). To confirm and update information obtained from these records, all patients were sent a questionnaire inquiring about the presence of a family history of IBD. Individuals identified as having other affected first-degree relatives were invited to participate and asked for permission to contact other affected and unaffected family members. Endoscopic, histological and radiological reports as well as clinical data were obtained on all affected individuals and these reports were reviewed for verification of diagnosis based upon standard criteria. Venous blood sampling was performed on affected individuals and their parents, and DNA was extracted using a salting out procedure. Ethics approval for this study was given by the University of Toronto Ethics Committee and written informed consent was obtained from all participants.

All of the LD analyses in this study were performed with father-mother-affected child (CD only) triads, where 0 or 1 of the parents was affected with CD. These triads either came from the multicase families used in the linkage stage of this study or were identified specifically for the purpose of the LD study. Specifically, for the microsatellite genotyping, 296 triads were genotyped: 95 of these triads were derived from families used in the original identification of the IBD5 locus (only one triad per family), and 201 were from newly collected families. For the SNP genotyping, 139 triads were genotyped: 18 were derived from families used in the original identification of the IBD5 locus, and 121 were from the newly collected families. Individuals affected by CD were identified by review of the clinical charts of all patients registered in the Mount Sinai Hospital Inflammatory Bowel Disease Centre patient database and from the Toronto Hospital for Sick Children IBD database. Written informed consent was obtained from all participants and ethics approval for this study was granted by the University of Toronto Ethics Committee.

Microsatellite Genotyping

Genomic DNA was extracted from peripheral blood lymphocytes from probands and family members from 163 Caucasian pedigrees. The genome-wide scan, with an average inter-marker spacing of 12 cM, was carried out using a modified version of the Cooperative Human Linkage Centre (CHLC) Screening Set/version 6.0 that also included Genethon markers. These 312 loci were amplified using fluorescently-labeled primers (Research Genetics Inc., Huntsville AL) in separate polymerase chain reactions, and the products were then multiplexed into panels by pooling before electrophoresis on ABI 377 sequencers (PE Applied Biosystems, Foster City, CA). Fluorescent genotyping gels were analyzed in an automated system developed at the Whitehead Institute/MIT Center for Genome Research. Further details of the genotyping system have previously been described (Rioux *et al.*, *Gastroenterology* 115:1062-1065 (1998)).

The region of suggestive linkage on chromosome 5 and the surrounding regions of poor information content were followed up with 34 additional microsatellite markers. Specifically, 34 markers were genotyped between markers D5S1470 and D5S1471, decreasing the average spacing between markers to approximately 3 cM in this 125 cM region. This higher density mapping was performed on the original samples and on additional 12 families, for a total of 175 pedigrees analyzed. These new families consisted of 16 CD affected sibpairs.

In the first phase of the microsatellite LD mapping, a total of 57 microsatellite markers were genotyped on 296 CD triads. Information regarding primer sequence, allele size range, and suggested amplification conditions for 55 of these genetic markers (all but IRF1p1 and CSF2p10) can be obtained from the Genethon (<http://www.genethon.fr/>), Marshfield (<http://research.marshfieldclinic.org/genetics/>), or Genome Database (<http://www.genethon.fr>) World Wide Web sites. The markers IRF1p1, CSF2p1, and the 8 markers used in the 2nd stage of LD mapping, were designed during the course of this study. Genotypes for all of these markers were obtained as described above.

SNP Discovery

In order to identify all SNPs in the *IBD5* critical region, a tiling path of overlapping PCR products was designed. Specifically, PCR assays were designed using Primer 3.0 to be approximately 700 bp in length, with 100 bp overlap with adjacent
5 assays. The -21 M13 forward and the -28 M13 reverse sequences were added to each of the forward and reverse PCR primers, respectively. These PCR primers were used to amplify 50 ng of genomic DNA from six CD patients, one unaffected family member, and one CEPH DNA as control. The PCR products were purified using the solid phase reversible immobilization (SPRI) method and then sequenced using the appropriate -21
10 M13 or -28 M13 DYEnamic Direct Cycle Sequencing kit (Amersham Pharmacia Biotech Ltd, Cleveland, OH). All sequencing reactions were run on ABI377 automated sequencers (PE Applied BioSystems, Foster City, CA); the gel files were processed using the BASS software, available on the Whitehead Institute/MIT Center for Genome Research FTP site. Sequences were base-called by the Phred program, and then the
15 forward and reverse reads were assembled by the Phrap program. All traces were visually inspected by at least two observers.

SNP genotyping

SNP genotyping was performed using length-multiplexed single-base extension (LM-SBE) as previously described. Briefly, PCR primers were designed as close as
20 possible to the SNPs identified in the current study, resulting in a product of a maximum length of 150 bp. Forward primers had T7 tails at their 5' ends and reverse primers had T3 tails at their 5' ends. These T7 and T3 tails were used for secondary amplification. Primer pairs were checked for homology to all amplicons and sorted into pools consisting of up to 50 primer pairs. Loci were subjected to two rounds of PCR
25 amplification. In the first round, 10 ng of genomic DNA was amplified using a pool of primer pairs (0.1 mM) and 2.5 units of Amplitaq Gold (Perkin Elmer). In the second round, a 3 mL aliquot of the primary amplification product was amplified with biotinylated-T7 and biotinylated-T3 primers. A 7 mL aliquot of this secondary

amplification product was purified from the unincorporated dNTPs using streptavidin-coated Dynabeads (Dyna). A multiplex SBE reaction was then carried out on the purified product using SNP-specific primers, JOE-ddATP (0.12 M), TAMRA-ddCTP (0.12 M), FAM-ddGTP (0.12 M), ROX-ddUTP (0.60 M; NEN DuPont) and

- 5 Thermosequenase (0.5 U; Amersham). Excess ddNTPs were removed from the SBE products using 96-well gel filtration blocks (Edge Biosystems) prior to electrophoresis on ABI 377 sequencers. The SBE gels were analyzed using a system developed at the Whitehead Institute/MIT Center for Genome Research as previously described.

Statistical analysis

- 10 Nonparametric multipoint linkage analysis of the data from the genome-wide scan and the higher density mapping on chromosome 5 was performed using the MAPMAKER/SIBS functions implemented in GENHUNTER 2.0. It is important to note that all sib pairs from sibships with more than 2 affecteds were counted but were conservatively downweighted by a factor of $2/n$ (where n = the number of affecteds).
- 15 Exclusion mapping was also performed with this software package, and a locus $8s > 2$ was considered excluded at a LOD score of -2.

- To establish appropriate thresholds for suggestive and significant genome-wide linkage for these particular datasets, simulations were performed by generating artificial genotype data with the identical family structures. These simulations matched the
- 20 datasets with respect to marker density, marker informativeness, the individuals genotyped, affected status, and the fraction of missing data.

- To assess the significance of the TDT results for each marker, permutation tests using the same genotype data were carried out. For each trio, chromosomes were randomly reassigned as transmitted or untransmitted to form a permuted dataset. The
- 25 number of permuted datasets with values as significant as that seen for the best single-marker and two-marker tests were tabulated. In order to quantify the extent of LD in the *IBD5* region, 3-marker haplotypes were examined using the TDT and P_{excess}
- (d). P_{excess} represents the strength of LD and is calculated by $(p_{\text{affected}} - p_{\text{normal}}) / (1$

- P_{normal}). In our study, the P_{affected} is calculated from the frequency of the haplotype among the transmitted parental chromosomes and P_{normal} is the frequency among untransmitted parental chromosomes.

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Table 1. Summary of the first stage of LD mapping using microsatellite markers.

Marker #	Marker Name	Source of marker ¹	Estimated Genetic Position ²	Distance to next marker	Previous linkage results (MLOD) ³	Allele	TDT results ⁴ X ²	pvalue
5	1 D5S1435	G	128.50	0.50	0.76	115	5.84	0.016
	2 AFMa113ye9	G	129.00	0.83		-	-	-
	3 D5S1505	M	129.83	0.00	0.79	-	-	-
	4 D5S1384	U	129.83	0.00		-	-	-
	5 D5S471	G	129.83	0.57	0.79	238	7.58	0.0059
10	6 D5S632	G	130.40	0.20		114	4.59	0.032
	7 D5S818	M	130.60	0.20		-	-	-
	8 D5S2502	M	130.80	0.10		-	-	-
	9 AFMB352XH5	G	130.90	0.04		-	-	-
	10 D5S1975	G	130.94	0.00		-	-	-
15	11 D5S622	G	130.94	1.86		-	-	-
	12 D5S2059	G	132.80	0.85		190	5.83	0.016
	13 D5S615	U	133.65	0.00	1.8	-	-	-
	14 D5S804	M	133.65	0.00	1.8	-	-	-
	15 D5S1495	M	133.65	0.00	1.8	382	4.00	0.045
	16 GATA68A03	M	133.65	0.35	2.2	-	-	-
20	17 D5S809	M	134.00	0.40	2.1	-	-	-
	18 D5S2120	G	134.40	0.20		-	-	-
	19 D5S642	G	134.60	0.65	2.6	-	-	-
	20 D5S2057	G	135.25	0.00	3.1	-	-	-
25	21 D5S2110	G	135.25	0.62	3.1	-	-	-
	22 IRF1p1	S	135.87	0.19		156	13.27	0.00027
	23 D5S1984	G	136.06	0.16		222	14.04	0.00018
	24 CSF2p10	S	136.22	0.58		307	4.00	0.045
	25 D5S2497	G	136.80	0.10	3.9	129	7.69	0.0055
	26 w2429/240wa7	G	136.90	0.10		-	-	-
30	27 w866/057vg5	G	137.00	0.10		-	-	-
	28 D5S1766	U	137.10	0.10	3.5	245	6.48	0.011
	29 D5S808	M	137.20	0.10	3.3	-	-	-
	30 D5S458	G	137.30	0.00	3.1	-	-	-
35	31 D5S396	G	137.30	0.09		-	-	-
	32 D5S2053	G	137.39	0.56	3.0	-	-	-
	33 D5S1995	G	137.95	0.69	2.8	-	-	-
	34 D5S2115	G	138.64	0.68	2.4	-	-	-
	35 IL9	M	139.32	0.01	2.0	-	-	-
40	36 D5S816	M	139.33	0.07	2.0	-	-	-
	37 D5S393	G	139.40	0.10	2.0	-	-	-
	38 D5S399	G	139.50	0.90	2.0	127	4.57	0.032
	39 D5S479	G	140.40	0.10		-	-	-
	40 AFM350yb1	G	140.50	0.10		-	-	-
45	41 D5S1983	G	140.60	0.12		116	4.55	0.033
	42 D5S476	G	140.72	0.00	1.7	-	-	-
	43 D5S500	G	140.72	0.28	1.7	211	4.15	0.042
	44 AFMB290YC9	G	141.00	0.82		-	-	-
	45 D5S414	G	141.82	0.98		-	-	-

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46	D5S2009	G	142.80	0.12		140	6.70	0.01
47	D5S658	G	142.92	0.00	2.0	-	-	-
48	D5S2116	G	142.92	1.08		-	-	-
49	D5S2011	G	144.00	0.06		-	-	-
50	D5S2119	G	144.06	0.00		-	-	-
51	D5S1979	G	144.06	1.15		-	-	-
52	D5S2017	G	145.21	2.19	2.2	91	5.40	0.02
53	D5S2859	M	147.40	0.09		-	-	-
54	D5S436	G	147.49	0.00	1.6	-	-	-
55	D5S207	M	147.49	0.00		-	-	-
56	D5S1480	M	147.49		1.6	-	-	-

¹ Abbreviations: G, Genethon; M, Marshfield; U, Utah; S, designed by authors from genomic sequence.

² Estimated from genetic (Genethon, Marshfield) and physical (data not shown) map information

³ Linkage data for the CD subgroup with early onset disease as seen in figure 1 and reference ###

15 ⁴ Results are shown only if pvalue < 0.05

Table 2. Summary of combined LD mapping information.

20

25

Marker #	Marker Name	Distance to next marker (kb)	LD mapping stage	Allele	T:U	TDI results X ²	pvalue
57	CAh14b	43	2	-		-	-
58	ATTh14c	167	2	-		-	-
59	IL4m2	164	2	214	1.8	4.74	0.029
60	GAh18a	21	2	193	1.4		0.018
22	IRF1p1	24	1	156	1.7	13.27	0.00027
61	CAh15a	130	2	373	1.5	7.73	0.0054
62	CAh17a	97	2	140	1.8	16.19	0.00005
						7	
23	D5S1984	163	1	222	1.8	14.04	0.00018
24	CSF2p10	178	1	307	1.4	4.00	0.045
63	CAh81b	85	2	-		-	-
64	CAh81c		2	-		-	-

Table 3

Legend:

EST: expressed sequence tag

gene: known gene

Predicted gene predicted from genomic sequence using the GENSCAN package

ins/del: insertion/deletion

genomic: derived from resequencing of entire genomic region (therefore includes genes, promoters, enhancers, etc.)

Notes:

1) "N" in sequence represents polymorphic base

2) details are provided where currently available

3) This list includes all polymorphisms: SNPs, repeats, and insertions/deletions

SEQ. ID NO:	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
1	CSF2_6610	c/t		Verified	gene	n/a	colony stimulating factor 2	aaacttcctgtgcaaccacagatatacacccttgaaagtttcaaaag
2	CSFenh_1492	g/t		Verified	gene	n/a	colony stimulating factor 2 (enhancer region)	atttcctcccctgtgataatgtctctcgtNataaggatcctggagtgactcaa gc
3	CSFenh_1580	g/t		Verified	gene	n/a	colony stimulating factor 2 (enhancer region)	acacgcataaggaaactcctcctccagagggtttcNcctgtctctgtaggaagg ggggcccccagaggg
4	CSFex4_6632	c/t		Verified	gene	n/a	colony stimulating factor 2	aaaggaaacttcctgtgcaaccacagatatacacccttgaaagtttcaaaaga
5	E4ex1_1	t/c		Verified	EST	n/a	n/a	cigggaacccaaacatcctgaggaaNagcigagaacacctaccaaggga
6	E4ex1_2	a/g		Verified	EST	n/a	n/a	agacagaaaattagcttagagatggagggtggcaNgaictctaaagctgt cccgcigcc
7	E4ex1_3	t/c		Verified	EST	n/a	n/a	atgggagggtggcac- gatctctaaagctgtccNgcgccatccaggagtgccctatgcataag
8	Facoex16_1	g/c		Verified	gene	n/a	Fatty acid CoA ligase	ggctactgaaagatccagacaggatNgaaggaggccctggacacgcat ggc

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
9	FaCoex1_1	t/c		Verified	gene	n/a	Fatty acid CoA ligase	accaggaggccctgctaccaccctgctaaNggctctaccaccaccggccttc
10	GENS010ex2_1	t/c		Verified	Predicted gene	n/a		agaagcagtagggcNactactaggtagcccca
11	GENS020ex1_1	a/g		Verified	Predicted gene	n/a		gggtgtgacagagggtgtNtggcaggactc
12	GENS020ex3_1	a/c		Verified	Predicted gene	n/a		ggcgccacNcaaacctctgtgcagctcc
13	GENS020ex3_2	t/g		Verified	Predicted gene	n/a		aggccagccctNttccttactatgtcct
14	GENS020ex3_3	a/g		Verified	Predicted gene	n/a		tagaagcagaagggtgtgtggcctcNctgtgtggacttctgccccactg
15	GENS021ex1_1	g/c		Verified	Predicted gene	n/a		cac
16	GENS021ex1_2	t/c		Verified	Predicted gene	n/a		tcatggcgggtgtgtgacctgagagaggNtcagatggaagaagccctg
17	GENS025ex1_1	a/g		Verified	Predicted gene	n/a		ggtaggaatgag
18	GENS026ex12_1	a/t		Verified	Predicted gene	n/a		aaggccctcatgtatcatgattNgtgtgtgtgtgtgtcct
19	GENS026ex3_1	t/c		Verified	Predicted gene	n/a		gctcaagccctggggagggaaggaagtggtgacccac
20	GENS026ex4_1	t/c		Verified	Predicted gene	n/a		cttcatgtagaaagagctagtagtactgtattNtataatgcttaccatgctcat
21	GENS026ex5_1	t/c		Verified	Predicted gene	n/a		atgaacaagcttc
22	GENS026ex5_2	a/g		Verified	Predicted gene	n/a		tccttctcacaactccttaagctaccNtgagagcaataggactcctgttaa
23	GENS026ex6_1	t/g		Verified	Predicted gene	n/a		gggtttgtgtatctaaataggNgacctcagccttaaacctcatct
24	GENS026ex6_2	t/c		Verified	Predicted gene	n/a		tggaaaaatcaattacccccgtattacNtgtgtgagagaatgaagcatt
25	GENS027ex2_1	a/g		Verified	Predicted gene	n/a		cagtaaatatgtaggcctatgtc
					Predicted gene	n/a		aatattattgtcttttaataaagtaNctctcgtcattgttgattcgtctatc
					Predicted gene	n/a		gta
					Predicted gene	n/a		tattattgtcttttaataaagtaNctcgtcattgttgattcgtctatcgtat
					Predicted gene	n/a		gcaatgctgtttttcttttagtatacaaaaNtgaatccttcttccctcaaaagctt
					Predicted gene	n/a		ga

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	Status Verification comment #3	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
26	GENS027pro_1	a/c		Verified	Predicted gene	n/a		ccccaccatctcgcggtggcggaagggaNaatggtatctttaataacaaa aagataat
27	GENS027pro_2	t/c		Verified	Predicted gene	n/a		atcttgaggcttatgaaccacacatatgttNgaaaacattgttgccctcctggc acaga
28	GENS02ex2_1	a/g		Verified	Predicted gene	n/a		ccatctatgtaggtaacNgaggcaaaagcaagggtaggagaga
29	GENS02ex3_1	g/c		Verified	Predicted gene	n/a		gggaggcagacattaggcaataatNacatggatctctgaaaaacatag ctctacga
30	GENS02ex4_1	t/g		Verified	Predicted gene	n/a		agaggaaatgggtggagttggcagNggggctgtctcggctctccccga
31	GENS030ex2_1	a/g		Verified	Predicted gene	n/a		ctggcttaggccaaaagaactggccaNgttacagttccacagagtaacccg
32	GENS030ex3_1	a/t		Verified	Predicted gene	n/a		agggtagtgagggtgactaggggaNctcggacacigagccccctgaagtgg gg
33	GENS030ex4_1	a/g		Verified	Predicted gene	n/a		gcggctgcagggggaggcacaagcNtggccaggcgccaagcggc
34	GENS031pro_1	t/c		Verified	Predicted gene	n/a		atgtctaccatggccaaactaatgtttga
35	GENS031ex1_1	t/c		Verified	Predicted gene	n/a		ctggtaaaacacaggctgccctggacaaaagcNggaacagaaatgaggct ccaggcgttgatt
36	GENS031pro_1	a/t		Verified	Predicted gene	n/a		ccacatttcttaatccagtcctatcattgNtggacattgggtgttccaagtcct tgc
37	GENS036ex1_1	t/g		Verified	Predicted gene	n/a		tccttcacaggacaggaattctgcacaaaNaacattcattagcttgcatgg taagcat
38	GENS036ex1_2	t/c		Verified	Predicted gene	n/a		aaatgggtactgtataccattacatctctgttttNgggggtgggtggcggggg ggga
39	GENS037ex1_1	a/g		Verified	Predicted gene	n/a		aatagggtgctgatttgcagtgacaatgagNcaattagtattatcaggagaa gctaacgaig
40	GENS037pro_1	a/c		Verified	Predicted gene	n/a		tgaactttagctctcttggtaaataaggaaaatNgctccaactactgtccaccc aagaaaac
41	GENS038ex3_1	a/c		Verified	Predicted gene	n/a		tatctgcgcctccctccacacagctgtcagNcttcatctaatgtgaaagac cagatgctcg
42	GENS038pro_1	t/c		Verified	Predicted gene	n/a		tccctccctgttctcgtcccgatctctgttNcatctatctatcgtggaggatt tctccaaact

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
43	GENS039ex4_1	a/g		Verified	Predicted gene	n/a		ccttctgaacatatttaataataacNaggaaaaacaataaattactcg ttggctga
44	GENS039ex7_1	a/g		Verified	Predicted gene	n/a		atgtgccttttctgctctccctcNtttcttagaagtcctccagaaaacc
45	GENS039ex7_2	t/g		Verified	Predicted gene	n/a		cgtgagtgccgctacttgccgtgtagcccccNctacggggccgtttcttaact gta
46	GENS03ex2_1	a/g		Verified	Predicted gene	n/a		ataatgcagaacaaaattagagaaaaactccNgtcagggtctccactcacc catggctggggct
47	GENS03ex6_1	a/g		Verified	Predicted gene	n/a		aaacaaaacatgccccgcagagtcaccNgggctggccatttgaaaaaga gtacatcag
48	GENS03ex6_2	a/t		Verified	Predicted gene	n/a		gggagggtctctggaaccagagagaccNglaggaggggactgcgggc aggagctgtg
49	GENS043ex1_1	a/g		Verified	Predicted gene	n/a		gcggcatctccatcttccaatgaactgagcNtgagcaatgaactgagtg acagtctcat
50	GENS043ex2_1	a/c		Verified	Predicted gene	n/a		tacttattcaattgcagttgtgaaaaaNtctgcaaatacglagccctc ccagttcaa
51	GENS044ex1_1	t/c		Verified	Predicted gene	n/a		cagtgtgctaggaaaagagatgtggattactgcNtctgtgcaatgataaag cagtaagtattccg
52	GENS044ex2_1	t/c		Verified	Predicted gene	n/a		tgtagtaaaaaacattcaaaaatctctctcNagctatcaagtatttgaatttg ctaaactggggctcattcttcctcagccNcattctgctaaatgccagatgcc ctgggaag
54	GENS044ex2_3	a/c		Verified	Predicted gene	n/a		tcgtcaatgccagatgccccgggaagNtcttctactgcccatttggaaggatg caga
55	GENS044ex2_4	t/c		Verified	Predicted gene	n/a		cctgggaagatcttctactgcccacNtggaaggatgcagaatgtgtgat ctgtcccatctccctataccatgtctgaNccccttagccataacatgtgatg gacagc
57	GENS045ex10_1	g/c		Verified	Predicted gene	n/a		aagctacacaagatgggcatttggcccttNaccaacatgctgttcttgactt cagcaaaccccatgcaaacattcagcatttcaNggctgaggccacacac agaagccatcag
58	GENS045ex10_2	t/c		Verified	Predicted gene	n/a		aaaccccatgcaaacattcagcatttccacNgtcaggggccacacacagaa gcc
59	GENS045ex10_3	a/g		Verified	Predicted gene	n/a		

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
60	GENS045ex10_4	g/c		Verified	Predicted gene	n/a		ggtagccacagatgtttctgtggclaccaacNgagaaaagccatctttaa acagc
61	GENS045ex10_5	t/c		Verified	Predicted gene	n/a		gccatctttaaacagcagagaaatctcactgttcNcctgtccactctctcct gtcaatcccaggac
62	GENS07ex1_1	t/c		Verified	Predicted gene	n/a		ccatctgagacctcatcagccagcgccttcactttccaNatcaccatcagcatt ctggttacaac
63	GENS09ex5_1	t/g		Verified	Predicted gene	n/a		ggggctgcgcgagcacctggggccNgggacgcagacccaa cagcactggccggggagcagaccccaaNacgacagcaggcagcgcc
64	GENS09ex5_2	a/g		Verified	Predicted gene	n/a		gagcg
65	GENS09pro_1	t/c		Verified	Predicted gene	n/a		tggaaagggccgacatggcaatgaatcta
66	IGR1000a_1	a/g		Verified	genomic	513		cccaggttggttttNgaactctggctt
67	IGR1002a_1	g/c		Verified	genomic	418		actgctggcgNgltgggtggt
68	IGR1002a_2	t/c		Verified	genomic	422		gctggcggggtgNgggtgctcacc
69	IGR1002a_3	g/c		Not yet verified	genomic	477		aggcaggtggatcacNaggtcaagga
70	IGR1002a_4	other/w+	Poly t	Verified	genomic	259		gtaaaatttaNtttttt
71	IGR1002a_5	a/t		Verified	genomic	405		ttagaaaaacNactgctggcg
72	IGR1003b_1	a/c		Verified	genomic	210		ctcagaaaaacaaaaaNaacaaaaagaaac
73	IGR1003b_2	other/w+	Poly t	Verified	genomic	1		taaaaatttaaNtttttttt
74	IGR1004a_2	other/w+	Poly a	Verified	genomic	395		aaaaaNaacaaacacttagag
75	IGR1006a_1	t/g		Verified	genomic	389		aactcctgacctaaNgatccgcctgctt
76	IGR1006a_2	ins/del		Verified	genomic	169		gtttttttNtttgagacagaa
77	IGR1007a_1	t/c		Verified	genomic	190		ttcctttaccatNctgtcctcatat
78	IGR1007a_2	t/c		Verified	genomic	196		ccatcctgtcNtcatatataaact
79	IGR1008a_1	other/w+	Poly t	Verified	genomic	605		tgggtgctctacNttttttt
80	IGR1008a_2	t/c		Verified	genomic	385		tattttgctcNgtggattctcct
81	IGR1009a_1	t/c		Verified	genomic	373		gtgctgggattaNaggtgtgaaccac
82	IGR1009a_2	t/c		Verified	genomic	389		agggtgaaccacgtNtccagccacttc
83	IGR1010a_1	other/w+	ca repeat	Verified	genomic	186		ttcattatgcacatNacacacacac
84	IGR1011a_1	g/c		Verified	genomic	207		ttcatccactgtgNacagtgtattt
85	IGR1012a_1	t/g		Verified	genomic	520		ggaaatcgcacaaaNaaacatttatta
86	IGR1012a_2	t/c		Verified	genomic	556		ggtaagcattgtcNtgcctgcctgt
87	IGR1013a_1	t/c		Verified	genomic	247		accattaccatctgctttNgggtgggtggcgcg

Table 3

SEQ. ID NO:	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
88	IGR1015a_1	a/c	a in ref. sequence	Verified	genomic	202		tccctcttgagtgctctcaNcggtctcgtgggtac
89	IGR1016a_1	a/g		Verified	genomic	300		cagccaccatcNtctagccgtgtt
90	IGR1016a_2	ins/del		Verified	genomic	420		atctgcttcNatgctttcccc
91	IGR1016a_3	t/g		Verified	genomic	103		ccctacaaccNatcgtcag
92	IGR1017a_1	t/c		Verified	genomic	537		aagggtgctgcagctccNaaggagtgttagaa
93	IGR1019a_1	t/c		Verified	genomic	366		gagcagcacatggNccaagttagagctaaag
94	IGR1020a_1	t/c		Verified	genomic	590		tccaccagccagaggtaactaNgctgttaatt
95	IGR1021a_1	t/g		Verified	genomic	237		gggtattagagaacaNgggattgagagctgc
96	IGR1021a_2	g/c		Verified	genomic	314		gcagattttgNtctglaaat
97	IGR1021a_3	t/c		Verified	genomic	411		agttcataatttaaNgtttttcagg
98	IGR1021a_4	ins/del	2 bp deletion	Verified	genomic	187		cttcttactctNtacctataccat
99	IGR1022a_1	g/c		Verified	genomic	402		aaccctctaaagatatcttNaaaggacttclaaaggaa
100	IGR1022a_2	g/c		Verified	genomic	522		glgcaaggccctaacgttttaNttgctctggtatcgca
101	IGR1022a_3	t/c		Verified	genomic	608		tctagctctggctgNtgcgtgctgctgcccag
102	IGR1023a_1	a/c		Verified	genomic	477		tttgtaaatagggaaatNgctccaactactgtc
103	IGR1025a_1	other/w+	ca repeat	Verified	genomic	557		ggagattttataNacacaca
104	IGR1026a_2	other/w+	Poly a	Verified	genomic	429		ccctatctcaNaaaaa
105	IGR1026a_3	ins/del		Verified	genomic	520		atgaaatgagatagctccagctaaaNgcccgaagag
106	IGR1027a_1	a/g		Verified	genomic	480		agagcaagctNaggagctc
107	IGR1027a_2	g/c	g on ref. sequence	Verified	genomic	497		gctctggacggcNagccccggaacc
108	IGR1029a_1	a/g		Verified	genomic	497		acaatlgagNcaattagttt
109	IGR1030a_1	a/g		Verified	genomic	554		agcactggggNacaatggt
110	IGR1031a_1	other/w+	Poly t	Verified	genomic	200		tcaggaatgacNttttttt
111	IGR1031a_3	a/c		Not yet verified	genomic	565		aagagctacNgtcttaccaa
112	IGR1032a_1	t/c		Verified	genomic	175		ccctaccccNagcagtga
113	IGR1032a_2	other/w+	Poly t	Verified	genomic	352		tatgaatttcNttttt
114	IGR1034a_1	a/g		Verified	genomic	293		tgcaatggcNcagctcagct
115	IGR1039a_1	t/g		Verified	genomic	462		ccttggcacNctactcagct
116	IGR1040a_1	t/c		Verified	genomic	188		cggccagaNgggcccctccc
117	IGR1040a_2	t/g		Verified	genomic	356		aggatttcaNgcaggaaagt
118	IGR1040a_3	a/c		Verified	genomic	633		agctgtcagNcttcatctaat

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
119	IGR1043a_1	a/g	g in ref. sequence	Verified	genomic	170		ggatctgcacNggaaggaatt
120	IGR1043a_2	a/g		Verified	genomic	377		gtactttgttNattaaataat
121	IGR1045a_2	t/c		Verified	genomic	200		ttgacaaaNtggccatga
122	IGR1045a_3	a/t		Verified	genomic	291		tagaagatttNaaaatgtaa
123	IGR1045a_4	t/c		Verified	genomic	99		cacacgcicNaatcaagccaccccaa
124	IGR1046a_1	t/g		Verified	genomic	301		gtcattggNtccccctcccc
125	IGR1046a_2	t/c		Verified	genomic	337		tctctgttNcatctatc
126	IGR1046a_3	t/c		Verified	genomic	572		tccatctNgttgaatg
127	IGR1047a_1	ins/del		Verified	genomic	253		agagcacaNacacatgga
128	IGR1050a_1	a/g		Verified	genomic	235		ctagatgaaggccataNgcagaagacattt
129	IGR1050a_2	a/t		Verified	genomic	558		gggtctgggtcccgNngtgcacagggg
130	IGR1052a_1	t/c		Verified	genomic	319		ccctcgtaaatatccttNcagccttaaacct
131	IGR1055a_1	a/g		Verified	genomic	566		atttaatacNaggaaaaacaat
132	IGR1056a_2	t/g		Verified	genomic	235		tattaccaggactccttggNgtccactgttttag
133	IGR1056a_3	t/c	this base is missing on ref. seq.	Verified	genomic	285		aacccttggctccaagtcNagcagccacagcttc
134	IGR1057a_1	t/c		Verified	genomic	271		ttcgaagtttcagttgaacNgtccctcgcgaaa
135	IGR1057a_2	a/g		Verified	genomic	390		gacaaagaggtcagcacNtagtagaacgc
136	IGR1060a_1	g/c		Verified	genomic	279		aaggagcggacitctactaaNgaatcctctgtaagg
137	IGR1060a_2	t/g		Verified	genomic	306		tgtaaagggcggccctatNtagtgcctggggagaat
138	IGR1063a_2	a/g		Verified	genomic	425		tctgtctctccctcNtttctctagaagtcctcc
139	IGR1064a_1	t/g		Verified	genomic	335		tggccgtgtgaccccNctacgggctgtttccta
140	IGR1066b_1	a/g		Verified	genomic	90		taccaaagggccgcctcNggcacttggcgcatgtg
141	IGR1068a_1	other/w+	poly T	Verified	genomic	141		ttcttaggtgtgNttttttttttt
142	IGR1070a_2	t/c		Verified	genomic	614		ttccattgtttcaNttggaatttatattttaatgt
143	IGR1070a_2	t/c		Verified	genomic	614		ttccattgtttcaNttggaatttatattttaatgt
144	IGR1070a_3	t/g		Verified	genomic	308		tctaactgtNtctaaactg
145	IGR1071b_1	t/c		Not yet verified	genomic	115		ttattccattgttttcaNttggaatttatatttta
146	IGR1072a_1	t/c		Verified	genomic	337		ctgacatatttttttaNttattagttattttttga
147	IGR1092a_1	a/c		Verified	genomic	241		aagcagagccaNacatacatctcac

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking sequence
148	IGR1095a_1	a/g		Verified	genomic	148		agaaggaggactNctggagccagg
149	IGR1095a_2	g/c		Verified	genomic	213		ttttctcggccaNcatagtccttatgca
150	IGR1098a_1	a/c		Verified	genomic	237		gcaagccagagNgacagggccacag
151	IGR1098a_2	g/c		Verified	genomic	294		ccctcttttgaatNcaaacctcgtc
152	IGR1099a_1	other/w+	Poly t	Verified	genomic	216		atgcattggcatgttcNttt
153	IGR1099b_2	a/g		Not yet verified	genomic	406		tagagacNgagtttcacc
154	IGR1099b_3	a/g		Not yet verified	genomic	270		ctggagNcaatggcag
155	IGR1100a_1	ins/del	deletion of 1 g t on ref. sequence	Verified	genomic	602		atgaaacctaacggNtctcagctctgttcta
156	IGR1100a_2	a/t		Verified	genomic	103		tgatttagaattttattNaaaaaaagtcaa
157	IGR1102a_1	t/c		Verified	genomic	605		ttttcttatNgcattttggct
158	IGR1102a_2	t/c		Verified	genomic	400		aattagccaggNgtagggagcgcca
159	IGR1102a_4	a/g		Not yet verified	genomic	119		ctgacattaccagNggaaaaaatggcgt
160	IGR1102a_6	other/w+	Poly a	Verified	genomic	549		cgagactccatctggNaaa
161	IGR1103a_1	other/w+	Poly a	Verified	genomic	78		aaaNgagtttcctcgtg
162	IGR1104a_1	g/c	g on ref. sequence	Verified	genomic	526		cagcttctaattgtgNttttattcctcag
163	IGR1105a_1	a/g	g in ref. sequence	Verified	genomic	383		ttaggttctttggaagcNggtttatgaactaat
164	IGR1107a_1	a/c	a in ref. sequence	Verified	genomic	402		aagattcaatgNaatcagtgacttgt
165	IGR1109a_1	a/g	a in ref. sequence	Verified	genomic	415		ggtagatgtgNtattacaaagatg
166	IGR1110a_2	ins/del		Verified	genomic	195		aaaaaNttattaccgg
167	IGR1111a_1	other/w+	Poly a	Verified	genomic	481		gagctagactcgtctcNaaa
168	IGR1111a_2	a/g	g in ref. sequence	Verified	genomic	318		tctactaaaNatacaaaa
169	IGR1111a_3	a/g	g in ref. sequence	Verified	genomic	325		atacaaNaattagcc
170	IGR1112a_1	t/c	c on ref. sequence	Verified	genomic	183		aaatacaaatagaNaacatacaaaa

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
171	IGR1113b_2	a/c		Not yet verified	genomic	293		tacctgaNgtgtgtctg
172	IGR1114a_1	t/c	c on ref. sequence	Verified	genomic	254		gtggctcacacNtgaatcccagcac
173	IGR1114a_2	a/g	a in ref. sequence	Verified	genomic	312		cccaggaagtcNaggctgcagtg
174	IGR1115a_1	other/w+	Poly a	Verified	genomic	465		gagccagactctgtcttNaaaaa
175	IGR1115a_2	a/c		Verified	genomic	307		ctctatctactaaaNatacaaaaattag
176	IGR1115a_3	t/c		Verified	genomic	322		atacaaaaattagcNgggtgtgtgtggg
177	IGR1115a_4	g/c		Not yet verified	genomic	438		gaatgaactccagcNtgggtgacagagcc
178	IGR1116a_1	t/c		Verified	genomic	625		gacttaaggtagcNctgaataaagccct
179	IGR1118a_1	a/g		Verified	genomic	192		gtatatttattagatNggglaatacatccaaatg
180	IGR1118a_2	other/w+	Poly a	Verified	genomic	47		ggcaaaaagagcgaactctgtctcaaaaaaN
181	IGR1118a_3	t/c		Verified	genomic	619		agcctgtgtgtctctaaNaagcctaaattgtctagaa
182	IGR1119a_1	a/g		Verified	genomic	190		ccaagctccctcatagNtccctatctgtctcag
183	IGR1120a_1	ins/del		Verified	genomic	258		ttttctttttttNctgagacagttttttc
184	IGR1126a_1	other/w+	Poly a	Verified	genomic	196		agagactccgtctcNaaaaa
185	IGR1142a_1	ins/del	deletion of 2 bp	Verified	genomic	526		ttttcgcaglaatacNtattaaaaatttagattc
186	IGR1142a_2	t/c		Verified	genomic	321		cagaacccctcatagcatgNtatcacigataaag
187	IGR1144a_1	t/c		Verified	genomic	435		catcaacaagggtcttaNagaattcctaagg
188	IGR1144a_2	a/g		Verified	genomic	611		aaatgagaaaaatctaNaatgaatctcgt
189	IGR1145a_1	a/g		Verified	genomic	338		tatcacttctcagtNataaagttcttaa
190	IGR1145a_2	g/c		Verified	genomic	463		aacaggtaftaattatcttcacattNcagtaataaagac
191	IGR1148a_1	other/w+	Poly T	Verified	genomic	304		tttttagagNtttttt
192	IGR1157b_1	t/c		Not yet verified	genomic	301		aagtgctggNatalacac
193	IGR1161a_1	t/c		Verified	genomic	221		cagtcctattttcaaaaNgagcaaacagaca
194	IGR1161a_2	t/c		Verified	genomic	662		aaactattttactaaaNagaagtcctccatta
195	IGR1169a_1	other/w+	Poly a	Verified	genomic	384		aaactctatcttNaaaaaaaaaaaaa
196	IGR1169a_2	a/c		Verified	genomic	454		tgtgtgcaNagtaagagaa
197	IGR1172a_1	t/c		Not yet verified	genomic	587		cctaacattatNttcaaaaaataa

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
198	IGR1173a_1	a/t		Not yet verified	genomic	517		agtttttNaaattttt
199	IGR1185a_1	t/c		Verified	genomic	516		aaaaattaNaaaaatagc
200	IGR1185a_2	t/c		Verified	genomic	576		aggctgaggNatgggaatc
201	IGR1186a_1	a/t		Verified	genomic	210		aacaagcttNictttaaac
202	IGR1186a_2	ins/del		Verified	genomic	423		ttttttNagctctgattc
203	IGR1193a_1	a/g		Verified	genomic	343		atgctagcNatgtaaaaaa
204	IGR1196a_1	t/c		Not yet verified	genomic	109		aaaaaaacaNaaggcact
205	IGR1196a_2	a/g		Not yet verified	genomic	202		gaagggtcaNacaggaaaag
206	IGR1196a_4	t/c		Verified	genomic	457		ggagcaaaaaNaaatgttta
207	IGR1199a_1	a/g		Verified	genomic	201		atatattccNagaaaatgcat
208	IGR1199a_2	t/g		Verified	genomic	214		aaatgcatcaNtaggcaattt
209	IGR1200a_1	other/w+	Poly a	Verified	genomic	516		gacgacctttNaaaaaaaaa
210	IGR1218a_1	t/c		Verified	genomic	469		ttttaataacNtgtaaaatgcc
211	IGR1218a_2	a/g		Verified	genomic	590		gcctgctggNtgagaggt
212	IGR1219a_1	t/c		Verified	genomic	129		gcttttaaaNttttct
213	IGR1219a_2	t/c		Verified	genomic	195		ctacaaagtNtattaaaggg
214	IGR1219a_3	a/t		Not yet verified	genomic	251		ttttgcttcaNagccttcctt
215	IGR1258a_1	other/w+	gt repeat	Verified	genomic	177		taaaactatataNgtgtgtgt
216	IGR1258a_2	t/c		Verified	genomic	436		tctgggagtaNtggcacaca
217	IGR1279a_1	g/c		Verified	genomic	223		accagtaattatttaaaaaatNaaagtaactaattgtt
218	IGR1279a_2	t/g		Verified	genomic	569		agccggggcgtgggtggcagNtgcctgtaatcccagct
219	IGR1286a_1	g/c		Verified	genomic	365		gttttgagaNagtcacact
220	IGR1319a_1	a/c		Not yet verified	genomic	200		taattttaaaggctctgNtccctgctctttc
221	IGR1350a_1	t/c		Verified	genomic	125		actctctcNtccccagg
222	IGR1353a_1	g/c		Verified	genomic	643		ctccaaaggaNctctgtctcc
223	IGR1353a_2	t/c		Verified	genomic	438		tggatggaNggacgaac
224	IGR1370a_2	ins/w+		Verified	genomic	172		taggggaggNcattccag
225	IGR1362a_1	t/c		Verified	genomic	434		caaggggaagNgcattccag
226	IGR1363a_1	a/g	G in ref sequence	Verified	genomic	382		gcagtgggNcaagtgtgg

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
227	IGR1364a_1	del/w+		Verified	genomic	147		gtttgttNgTTTTtgag
228	IGR1365a_1	t/c	ins/w+	Verified	genomic	160		actgggatgNtcctaaactg
229	IGR1365a_2	ins/w+		Verified	genomic	211		gacttttNaatagagat
230	IGR1366a_1	a/g		Verified	genomic	371		caagacagtgNataaatagc
231	IGR1367a_1	a/g		Verified	genomic	73		aaagaaaaNlcagaattt
232	IGR1367a_2	del/w+		Not yet verified	genomic	425		cctccttccccNcttctc
233	IGR1369a_1	a/c		Verified	genomic	44		tcaaaagagaNcaatgatga
234	IGR1369a_2	a/c		Verified	genomic	91		aaagtactaNtatgaaat
235	IGR1370a_2	del/w+		Not yet verified	genomic	350		tatatataNacacacatc
236	IGR1370a_3	t/g		Verified	genomic	241		gaagaaaaNagtgcagtg
237	IGR1371a_1	t/c		Verified	genomic	72		aaaatatgcNlcaggagtg
238	IGR1371a_2	a/g		Verified	genomic	231		aaaaaaagNccaacagaaa
239	IGR1372a_1	other/w+	poly t	Verified	genomic	298		tttttttNaggagagtg
240	IGR1372a_2	t/c		Verified	genomic	323		ttctgtgctcNggctggagt
241	IGR1373a_1	t/c		Verified	genomic	338		aacttagaaNtctccagg
242	IGR1375a_1	t/c		Verified	genomic	96		aggaattgaaNttaataga
243	IGR1376a_1	a/t	A in ref sequence	Verified	genomic	462		cacttgtnlgattaat
244	IGR1376a_2	del/w+		Verified	genomic	79		gcaagaagcNcaacaaacc
245	IGR1380a_1	other/w+	poly t	Verified	genomic	573		agtcaccaacNttttt
246	IGR1380a_2	a/g		Verified	genomic	155		ttaatatgatNaaatgctcaa
247	IGR2001b_1	a/c		Verified	genomic	148		cccccaaaagNccgagaagcct
248	IGR2002a_1	a/c		Verified	genomic	357		aaaatcgagatgaaggNttgagcatttcagaga
249	IGR2003a_1	a/g		Verified	genomic	234		ttgcagtgagccNagatcacgtcact
250	IGR2004a_1	ins/del	deletion of 14 bp	Verified	genomic	576		tagagttgttcccNagagttgttccca
251	IGR2006a_1	t/c		Verified	genomic	122		ctttagtttcatcttNcctactgcca
252	IGR2006a_2	a/g		Verified	genomic	380		ctggctccNaattaataag
253	IGR2007a_1	other/w+	Poly a	Verified	genomic	459		taaagtaagaatcccctaaggttNaaaaaataaaag
254	IGR2008a_1	t/c		Verified	genomic	646		ttactctgcaggagctNtagggagatgaaggaagcc
255	IGR2008a_2	g/c		Verified	genomic	596		cccggaggaggagcgtNggtagaggaaatgacac

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
256	IGR2009a_1	ins/del	deletion of "c"	Verified	genomic	270		agagtaagtagggNccttaccagagcat
257	IGR2010a_2	t/c		Verified	genomic	359		aggcttctgccNcttcacttcccca
258	IGR2010a_3	a/g		Verified	genomic	233		ggtagggctactNttatttatggtt
259	IGR2010a_4	a/g		Verified	genomic	113		ccctggcactatNacccctgaacggcg
260	IGR2010a_6	a/g		Verified	genomic	329		agcacacggggcaNggtaggtcttcgcc
261	IGR2011a_2	a/g		Verified	genomic	43		ggcgatcacctcNcctgcgttcggg
262	IGR2011a_3	a/g		Verified	genomic	153		acaggctggggccNggggcgctgggc
263	IGR2011b_1	g/c		Verified	genomic	396		agacgtgccccgagccccgcgaaNcgaggccaccggagccgtgc cc
264	IGR2011b_2	t/c		Not yet verified	genomic	500		CCACTCGGAGTCGGCTNCGGGGGCCCTCACTGCA
265	IGR2013a_1	g/c		Verified	genomic	431		GCCCC
266	IGR2015a_1	t/g		Verified	genomic	443		aaaggattgaattttgagNgaagaagt
267	IGR2016a_1	a/t		Verified	genomic	366		ctgcagtagtctctgTgggNtagatcttaaatgtc
268	IGR2016a_2	a/g		Verified	genomic	120		ggaagaagttcttacttccNtgtgggtgctta
269	IGR2017a_1	t/c		Verified	genomic	412		actcataattNtactgtgtccc
270	IGR2018a_1	t/c		Verified	genomic	245		ggtccctgagctcccNgagacaacatgcagaattactg
271	IGR2020a_1	a/c		Verified	genomic	568		gtcagccacccattNagtaactgttctctgtg
272	IGR2020a_15	t/g		Verified	genomic	408		gagagagaaaagatgNtcagaactccaccctggcac
273	IGR2020a_2	a/g		Verified	genomic	379		tcctccgactNgcacatccagt
274	IGR2020a_3	t/c		Verified	genomic	362		ccccagcacgtgccNtgtctgcagcagcacctccc
275	IGR2020a_4	a/g		Verified	genomic	301		accgtggctctgcgtNccccagcacgtcgcc
276	IGR2020a_5	a/g		Verified	genomic	210		gcagggtgtgtcggNggcgctcgtctgtgcaactaa
277	IGR2020a_9	a/g		Verified	genomic	194		caggctggcaggNgacccacacaggtcagtggaatgactc
278	IGR2021a_1	ins/del		Not yet verified	genomic	233		actccaggtagctgNtccaggctcggc
279	IGR2021a_2	a/g		Verified	genomic	147		ggccaggggtgcattttgNgggtgttctctctc
280	IGR2021a_3	t/g		Verified	genomic	197		ccatagggggagggcaagcgacNgggacactaggaaggca
281	IGR2021a_4	other/w+	gt repeat	Verified	genomic	394		ctgcagtagctgggggtgNtgagagggagggagggg
282	IGR2021a_5	ins/del	deletion of 16 bp	Verified	genomic	277		gtgtgNcagagagacagagagacagagagagag
283	IGR2022a_1	t/c		Verified	genomic	612		gcccagcatctgagggNtaggggtglaatcggca
284	IGR2022a_2	t/c		Verified	genomic	439		aggctcaggagttNgagaccagcctgactaacatggtgaaa
285	IGR2022a_3	t/c		Verified	genomic	190		aatcagccttaggatcNgttaatatgatgtgcttt
286	IGR2022a_4	a/g		Verified	genomic	248		ctgtgtcacctggctgNttgattgtcccaagtc
				Verified	genomic			ggaaagccaccatNggaaaggaaggcagg

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
287	IGR2024a_1	t/g		Verified	genomic	163		gccaaagggtgatactggctNagaggagctggctca
288	IGR2024a_2	a/g		Verified	genomic	461		atggagaaagcttggggcaggNccaggagcagg
289	IGR2024a_3	t/g		Verified	genomic	517		cacattgaattagctacNcgtgccatgccttaagg
290	IGR2024a_7	a/g		Verified	genomic	468		gggcaggccaggNgcagggcggtaaa
291	IGR2025a_1	t/c		Verified	genomic	139		ccgatgccaccgtcccNtaccctcatacaac
292	IGR2025a_2	a/g		Verified	genomic	141		ctgatccaccgtccctNccctcatacaacctctt
293	IGR2025a_5	a/g		Verified	genomic	270		ttgccctccatccaNgccattccctgt
294	IGR2025a_6	a/g		Verified	genomic	377		aagctggactctgtNggcccccaac
295	IGR2026a_1	ins/del	deletion of "c"	Verified	genomic	244		cacaaagaactaccccNtttcagctgagccc
296	IGR2026a_2	a/g		Verified	genomic	314		gtgggtctctggggcNatgctccctcagcctc
297	IGR2026a_3	ins/del	ins/del "a"	Verified	genomic	611		tcatgtlgaacacataNgacgtgtgtaaatgta
298	IGR2027a_1	ins/del	ins/del "g"	Verified	genomic	166		aaagtaaatgtttataaNgggtgtggccttttagaga
299	IGR2027a_2	a/g		Verified	genomic	291		gaacagggaatgcatcNttataaaatccttgc
300	IGR2027a_3	a/c		Verified	genomic	309		ttataaatccttgcgNcaggcgggtggctcacacctg
301	IGR2027a_4	t/c		Verified	genomic	386		tcactgaggtcaggagtNagaccagcctggtgaaa
302	IGR2027a_5	other/w+	Poly a	Verified	genomic	562		actcagccgggcaccNaaaaa
303	IGR2029a_1	a/g		Verified	genomic	112		tgaaccgggagatgNaggtgcagtgagct
304	IGR2029a_2	t/c		Verified	genomic	166		tcagccctgggtgacaagNagacttgttctcaaa
305	IGR2029a_3	other/w+	Poly a	Verified	genomic	180		tgtctcaaaaaaaataatcctttg
306	IGR2030a_1	t/g		Verified	genomic	539		gaagggtggatatgtcNtttcgtctcct
307	IGR2031a_1	t/g		Verified	genomic	415		gatcgtgtgagtgccaggNggactcctgtcgggta
308	IGR2031a_3	t/g		Verified	genomic	40		tgtggatatgtcNtttcgtctcct
309	IGR2031a_4	a/g		Verified	genomic	227		ctcagtcacagaaaccNtatgtactgtgac
310	IGR2031a_5	t/g		Verified	genomic	232		ctcagtcacagaaaccatagtNacttgaccccgctcact
311	IGR2032a_1	ins/del		Verified	genomic	126		tcttactaaaaaNaactaaccaggcgtgtg
312	IGR2032a_2	a/g		Verified	genomic	356		ggaacagaggNatagacagga
313	IGR2032a_3	other/w+	Poly a	Verified	genomic	278		agactctctcNaaaaa
314	IGR2033a_1	t/c		Verified	genomic	587		atcatttaaggNctgacagtgctctg
315	IGR2034a_1	t/g		Verified	genomic	441		gaagctaataNgaaccatc
316	IGR2036a_1	g/c		Verified	genomic	356		acctcaaaagtNtggctggata
317	IGR2036a_2	a/g		Verified	genomic	183		gtaagacacacNgcctgcagag
318	IGR2037a_1	ins/del	ct repeat	Verified	genomic	534		aagacaacctagctNcgttctgtctttaa
319	IGR2038a_1	ins/del	aaac repeat	Verified	genomic	532		tgagttctacacagtggtNaaacaaaca
320	IGR2039a_1	ins/del		Verified	genomic	394		tgtgtgctNgtggat
321	IGR2041a_1	a/g		Verified	genomic	331		cacgtataaagccaccctacNataaccacc

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
322	IGR2042a_2	t/c		Verified	genomic	270		gagggccaaaggcttgcctgccNcctcgccct
323	IGR2043a_1	a/g		Not yet verified	genomic	334		tcgtatagtgccNggaacatcctgact
324	IGR2047a_1	other/w+	Poly t	Verified	genomic	225		tgtagggctttgcNttttt
325	IGR2049a_2	t/c	t on ref. sequence	Verified	genomic	332		gaccttcgtctacaNgtacataacaatagcata
326	IGR2051a_1	t/c	t on ref. sequence	Verified	genomic	470		ggcagggNtgtctggccaaggaccagtcc
327	IGR2051a_2	a/g		Verified	genomic	605		acacttatNtaactgtcacccctggcccatt
328	IGR2052a_1	t/c		Verified	genomic	290		gctattttctcNttgtattctgcagtgaccaagg
329	IGR2052a_2	a/g		Verified	genomic	106		ttgacaaacacttatNtaactgtcacc
330	IGR2053a_1	a/g		Verified	genomic	225		cattcaactgtctgtctcNgggctagagaaga
331	IGR2053a_2	a/c		Verified	genomic	369		cactgcgtctgcagtgacNcctgcttccccctaagt
332	IGR2053a_3	t/c		Verified	genomic	544		gtgacctattggatcttctcaNggcactgaggatat
333	IGR2054a_1	t/c		Verified	genomic	196		caagaggggaatggagctcttNgcagaggggctg
334	IGR2054a_2	ins/del	ins/del 6 bp	Verified	genomic	591		cttctgctctctctgNccctctgcctc
335	IGR2055a_1	t/g		Verified	genomic	609		gagtgtggttgagaagaNtctgaggagtgggac
336	IGR2056a_1	a/g		Verified	genomic	153		ttttaaagactagtcNctgggctcggt
337	IGR2056a_2	a/c		Verified	genomic	364		gagaaatgggtgaacccggaggNagagcttgcat
338	IGR2056a_3	other/w+	Poly a	Verified	genomic	481		aagcgagactccatctcNaataaaaaaacaaaacaa
339	IGR2056a_4	g/c		Verified	genomic	432		gagctgcagtgagctgaNatcgccgacctgact
340	IGR2057a_1	a/g		Verified	genomic	421		gaaglgaaaaccataaNcaagggtctacaga
341	IGR2060a_1	g/c		Verified	genomic	514		ttgcaacctNgcaaaaggtaa
342	IGR2061a_3	t/c		Verified	genomic	236		catacacaagaaNgagttcatttacitg
343	IGR2062a_1	ins/del	caaaa repeat c on ref. sequence	Verified	genomic	195		aaaaaaaaaaaaaacaaaaaNaacatgcatgcc
344	IGR2063b_1	g/c		Verified	genomic	218		ggcaataatNacatggatctc
345	IGR2063b_2	t/g	t on ref. sequence	Verified	genomic	369		agttggcagNggggctggttc
346	IGR2064a_1	a/c		Verified	genomic	364		aaactgtgattNcagtttcattt
347	IGR2064a_2	a/g		Verified	genomic	508		ccctcagagggcNgggtactggact
348	IGR2066a_1	t/c		Verified	genomic	459		cttcatctccctgcoccaaNGaagctggtggtgccc
349	IGR2067a_1	a/g		Verified	genomic	163		agccactactgggcNgtcagctc
350	IGR2067a_2	a/g		Verified	genomic	243		cacacttctccacNagaaataaagcaagca
351	IGR2067a_3	t/c		Verified	genomic	266		agcaagcagctgttNctctcttgggcc

Table 3

SEQ. ID NO.	Polymorphism Name	Polypeptide	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
352	IGR2067a_4	a/g		Verified	genomic	485		agcctgagcctNgccagccagac
353	IGR2068a_1	other/w+	ca repeat	Verified	genomic	354		acacacacacaNtttttgagagag
354	IGR2068a_2	g/c		Verified	genomic	70		atgttagtgtgagaaNgtgagaggtactcg
355	IGR2069a_1	a/g	g in ref. sequence	Verified	genomic	394		ttatgttcattgtacNtttcaccatatttt
356	IGR2069a_2	t/c		Verified	genomic	425		atccactctcNgtcatggacatctg
357	IGR2070a_1	t/c		Verified	genomic	551		tctaaagaaaaaagagcNgtgaattcttgac
358	IGR2071a_1	t/g	g on ref. sequence	Verified	genomic	165		gctctgtgccaggcaggggNctccgaggtgagtg
359	IGR2071a_2	a/t	a on ref. sequence	Verified	genomic	171		ccaggcagggggctccgNggtagtggtggcct
360	IGR2071a_3	a/g	a in ref. sequence	Verified	genomic	365		agagaagggaactggcNgtgtggctgggctgtg
361	IGR2072a_1	a/g	a in ref. sequence	Verified	genomic	312		gcaggctcagtggaaggagaggNgtctcttatg
362	IGR2072a_2	t/c	t on ref. sequence	Verified	genomic	408		atgggaactctctctaNactgtggagggcgtg
363	IGR2073a_1	a/c	a in ref. sequence	Verified	genomic	94		agtcatggcactaNatggagcccaggg
364	IGR2073a_2	a/g	a in ref. sequence	Verified	genomic	313		caccaggaggttcagcNcccactgtgg
365	IGR2073a_3	t/c	c in ref. sequence	Verified	genomic	379		gcataccagcgccNggccagtggtcc
366	IGR2074a_1	ins/del		Verified	genomic	239		gagtaaggggtcNaggaggggggggtggc
367	IGR2076a_1	t/c	t on ref. sequence	Verified	genomic	184		gaacatactcataNccatgcttcccc
368	IGR2076a_2	other/w+	Poly t	Verified	genomic	647		tacactatggttggtgcNttttttt
369	IGR2077a_1	other/w+	Poly t	Verified	genomic	148		tatggttggtgcNttttttttt
370	IGR2078a_1	a/g	g in ref. sequence	Verified	genomic	197		gcaggggtgggagaaNgccagactcaggggtg
371	IGR2078a_2	ins/del	ins/del "c"	Verified	genomic	67		ggccagccccccccNggaggtggat
372	IGR2079a_1	ins/del	Poly a	Verified	genomic	345		gtaaaaaaaNccctacaggtaaaag
373	IGR2079a_2	t/c	t on ref. sequence	Verified	genomic	582		cccccatgtgccaNgtcaacctccctgtc

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
374	IGR2081a_1	a/g	a in ref. sequence	Verified	genomic	140		ccagcaggaacaNatgcaca
375	IGR2081a_2	a/t	t in ref. sequence	Verified	genomic	315		gaaccagagagaccNgtaggagggg
376	IGR2081a_3	a/g	a in ref. sequence	Verified	genomic	622		gccggcagagtcaccNgggctggcc
377	IGR2083a_1	t/c		Verified	genomic	372		aaatggggccaggNgcgggtgctca
378	IGR2083a_2	ins/del		Not yet verified	genomic	199		ccgtcttaaaaaaaaaNNNgcctgggtggtg
379	IGR2083a_3	a/g	g in ref. sequence	Verified	genomic	572		aattgctgaaccccNggaggcagaggtt
380	IGR2084a_2	ins/del	ccaa repeat	Verified	genomic	166		ccaaccaaccaNccaaatggtattactctc
381	IGR2085a_1	t/c	c on ref. sequence	Verified	genomic	131		cacttaccctgcccNgccccaccc
382	IGR2085a_2	a/g	a on ref. sequence	Verified	genomic	249		tcttcttgaacctNtgtgattct
383	IGR2085a_3	a/g	a on ref. sequence	Verified	genomic	437		tgtcaacagtcaccaNctgagcccagcc
384	IGR2085a_4	a/g		Verified	genomic	368		ctgaggctgctcNtaagagggtccaatga
385	IGR2085a_5	t/c		Verified	genomic	538		ttattccagtcaccNtagtcattccagtc
386	IGR2087a_3	other	gaa repeat	Not yet verified	genomic	193		aggaagaagaagaaNcaagaggaagagga
387	IGR2087a_4	other/w+	Poly a	Verified	genomic	504		gaaagccaaaataaaaaaaNicaacagaa
388	IGR2090a_1	a/c	a in ref. sequence	Verified	genomic	219		agtcaggctgtctcgccNgciaaaagagcc
389	IGR2090a_2	t/c		Verified	genomic	360		tgcctgtggggctcNagcgttaccgccg
390	IGR2090a_3	t/c	c on ref. sequence	Verified	genomic	444		ttcacccattgttctcNctattcccttt
391	IGR2090a_4	other/w+	Poly t	Verified	genomic	532		acttacctgtgaaatgcactgNttttttt
392	IGR2091a_1	t/g		Verified	genomic	581		taatgacattccctgtgtaNgaatgtgccaatgtgga
393	IGR2091a_3	a/c	a in ref. sequence	Not yet verified	genomic	391		galcacattaNttgcctgagtt
394	IGR2091a_4	t/c		Not yet verified	genomic	404		ttgcctgagttcNcaagttggttaagaga

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
395	IGR2091a_5	ins/del		Not yet verified	genomic	547		tcctcatcaataaattattNNNctcatcatt
396	IGR2092a_2	ins/del	Poly a	Verified	genomic	435		aaaaaaaaaaaaNggccaggcgcg
397	IGR2093a_1	ins/del	Poly a	Verified	genomic	229		aaaaaaaaaaNgccctagacccctcg
398	IGR2093a_2	a/g	g in ref. sequence	Not yet verified	genomic	123		ttggaggcctgaggcNgaagaatcgct
399	IGR2093a_3	t/c		Verified	genomic	181		agattgcccactgNgctcagctct
400	IGR2093a_4	a/g	g in ref. sequence	Verified	genomic	318		gggagaccggaggaggNtagggaagtg
401	IGR2095a_1	t/c	c on ref. sequence	Verified	genomic	421		caacagccggcagNgagggcctgtct
402	IGR2096a_1	a/c	c in ref. sequence	Verified	genomic	112		actagagggttttttaNagagaagtgcattgat
403	IGR2096a_2	a/g		Verified	genomic	498		taaggaatacgggttttgNacglaagtgtagatgct
404	IGR2097a_1	a/c		Not yet verified	genomic	58		caggtggaanltgtaatactggggagag
405	IGR2097a_2	ins/del	Poly a	Verified	genomic	463		aagactctgtctcNaaaaa
406	IGR2101a_1	t/c		Not yet verified	genomic	283		cccagaatagagaccacNtccatctcctt
407	IGR2102a_1	g/c	g on ref. sequence	Verified	genomic	166		gaacttagatttgcgNcccttagcattcaac
408	IGR2102a_2	t/g	g on ref. sequence	Verified	genomic	223		caatgcattgatcctNctgagccctcagc
409	IGR2105a_1	a/t		Not yet verified	genomic	493		ttgatactcagtaNgtacagcttatt
410	IGR2106a_1	other/w+	ct repeat	Verified	genomic	137		caggcaacaaaaNtctccctccct
411	IGR2107a_1	t/c	c on ref. sequence	Verified	genomic	300		ccttgcttcaaNtgcctcagctctatc
412	IGR2107a_2	t/c	t on ref. sequence	Verified	genomic	564		ccaaaggctNcaggctctggc
413	IGR2108a_1	ins/del		Verified	genomic	360		ccattccctgagcNcagggtgccttct
414	IGR2109a_1	a/g		Verified	genomic	400		ggccaggctggtctcNgtctagactcaagtg
415	IGR2110a_1	t/c		Not yet verified	genomic	286		tgtttgagacagggtcttgNtctgctccaggatgg
416	IGR2110a_2	other/w+	Poly t	Verified	genomic	420		atgccagctaNttttt

Table 3

SEQ. ID NO.	Polymorphism Name	Polymorphism type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
417	IGR2111a_1	a/g	c on ref. sequence	Verified	genomic	55		ccacgcacccggccaNttttattgttttaaa
418	IGR2111a_3	t/c		Verified	genomic	516		ttgccaacatttggtatNacagtcctcaatttt
419	IGR2112a_1	other/w+	Poly t	Verified	genomic	285		ttttttttNcigagacagagtcgcgt
420	IGR2114a_1	other/w+	Poly a	Verified	genomic	3331		caattgacttccctNaaaaa
421	IGR2117a_1	a/g		Verified	genomic	355		aagggtgacctagNgcacacactccctccc
422	IGR2121a_1	other/w+	Poly a	Verified	genomic	609		aataaagtgacttNaaaaaataaaaaa
423	IGR2121a_2	a/t		Verified	genomic	117		gaggccctgacagNttgaaggggtg
424	IGR2121a_3	t/c		Verified	genomic	815		cctcggggtNttccaaatca
425	IGR2123a_1	a/g	g in ref. sequence	Verified	genomic	230		ttgccagaacacNgggtcagagagcaagag
426	IGR2125a_1	other/w+	Poly a	Verified	genomic	546		agagtgagactctgtctcaaaaaataaaaaa
427	IGR2126a_1	a/g		Verified	genomic	364		cttcatactacttNgaataccatat
428	IGR2126a_2	other/w+	Poly a	Verified	genomic	47		gagactctgtctcNaaaaa
429	IGR2131a_1	other/w+	Poly a	Verified	genomic	249		aaaaaataaaaaNgaacctctgtcgta
430	IGR2134a_1	a/g	a on ref. sequence	Verified	genomic	339		acttcagattaataNgtcttaaccat
431	IGR2136a_2	t/c		Verified	genomic	444		tgctagctccatttgagNaggacctt
432	IGR2138a_1	a/g		Verified	genomic	375		atgatttgcNtcaaaagcag
433	IGR2144a_1	t/g		Not yet verified	genomic	384		tcagtlaccacatctgtNttccatgctctt
434	IGR2144a_2	other/w+	Poly a	Verified	genomic	463		acagaggtaaaagtggttttgaaagcNaaaaa
435	IGR2144a_3	a/t		Verified	genomic	127		ctagcctaNggtctaggccc
436	IGR2144a_4	t/c		Verified	genomic	137		ggtctaggcNctcctgcctg
437	IGR2144a_5	a/g		Verified	genomic	166		ggaatcattacNtatcaaatca
438	IGR2147a_1	a/g		Not yet verified	genomic	354		accatggatgcNtagctgagttcctg
439	IGR2148a_1	t/c		Verified	genomic	253		acagttgtccctNagcatcttcgagga
440	IGR2148a_2	other/w+	caaaaa repeat	Not yet verified	genomic	619		gagactctatNaaaaaataaaaaataaaaaa
441	IGR2150a_1	g/c		Verified	genomic	90		aaactcaccacNactgaaatctggtta
442	IGR2150a_2	t/g		Not yet verified	genomic	336		ccctgggggctctaNttttgggttac
443	IGR2150a_3	t/g		Verified	genomic	558		gaaagataNaaataataataaa
444	IGR2151a_1	other/w+	Poly a	Verified	genomic	202		aaaaaNtcataccaatagtcactataa

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
445	IGR2151a_2	a/g		Verified	genomic	566		catctgcaNcccagcttc
446	IGR2153a_1	a/g	a on ref. sequence	Verified	genomic	423		cagaacaaattagagaaaaactccNgtcaggctctccac
447	IGR2154a_1	a/g		Not yet verified	genomic	389		acaacaacgggtaNatatttttaggtctc
448	IGR2155a_1	a/c		Not yet verified	genomic	398		attattagtcNaataatcacc
449	IGR2155a_2	a/g		Not yet verified	genomic	619		aaggcggggtNcagtggtctcac
450	IGR2156a_1	a/c		Not yet verified	genomic	176		ctgaggcaggctggatcatNtgaggctcagg
451	IGR2157a_1	a/c		Verified	genomic	254		tggaaagagacatgcatNcaaaccatatc
452	IGR2159a_1	other/w+	Poly t	Verified	genomic	411		tttttttttNccgtgaacag
453	IGR2160a_1	other/w+	ca repeat	Verified	genomic	601		acaggcgcgcNcacacacacacaca
454	IGR2160a_2	a/g		Not yet verified	genomic	213		taaaaaattattcgNgagaatttttagaa
455	IGR2160a_3	a/g		Not yet verified	genomic	287		ccaagtacctggNctgtactgagagatga
456	IGR2162a_1	a/c		Not yet verified	genomic	350		acaaacaaacaaNcaaaccttatt
457	IGR2162a_3	t/c		Not yet verified	genomic	450		aaatatagNcaaaatact
458	IGR2164a_1	a/t		Not yet verified	genomic	557		tcttgccaacNtggtgaaacccc
459	IGR2165a_1	ins/del	Poly a	Verified	genomic	473		ggaaaaaaaNcacacatgat
460	IGR2165a_2	ins/del		Not yet verified	genomic	271		ataaaaaaaaNgattttattgt
461	IGR2166a_2	a/c		Not yet verified	genomic	323		agtttcNgtttagaaag
462	IGR2167a_1	t/c	c on ref. sequence	Verified	genomic	324		acttaagagaNcaaaataatttt
463	IGR2167a_2	a/t		Not yet verified	genomic	207		ttttaaaacttNtaaaggaat
464	IGR2168a_1	ins/del		Not yet verified	genomic	341		tgtttctttttttcttcttNtttttttagacggag

Table 3

Seq. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
465	IGR2175a_1	a/g	g in ref. sequence	Verified	genomic	310		tgggccaaaaatctcNtctgacttccagtg
466	IGR2175a_2	a/g	g in ref. sequence	Verified	genomic	526		tccaaggctcacatNgttactatgtatgtt
467	IGR2176a_1	a/g	g in ref. sequence	Verified	genomic	119		gaagcaagacgctcNggaacaciggaac
468	IGR2176a_2	a/g		Not yet verified	genomic	399		aaccatctgttggctcNtgaggctctctgtat
469	IGR2177a_1	a/c	c in ref. sequence	Verified	genomic	325		tgtatgacacgcaacNcagctgaagaatgat
470	IGR2178a_1	a/g		Verified	genomic	138		ccatctaaataactacaagatgcNttgacgtataaga
471	IGR2179a_1	a/g		Verified	genomic	284		aaagtcacaaaaatcNaaaggagatgagca
472	IGR2179a_2	other/w++	Poly t	Verified	genomic	371		tctgggaaaaaggaaagtcNtttttttttt
473	IGR2179a_3	ins/del		Not yet verified	genomic	470		taatctctgcctccaggNtcaagtgattctct
474	IGR2180a_1	a/g		Not yet verified	genomic	65		glattttttagtagagacNgggttccttatgtt
475	IGR2180a_2	g/c		Not yet verified	genomic	383		tcaccagcaacctgttNtgagtgaatc
476	IGR2181a_1	other/w++	Poly t	Verified	genomic	260		aaaaagttttttttNctaccacaaatgtacag
477	IGR2181a_2	t/c		Verified	genomic	416		attacattataattacaNgcaigatc
478	IGR2181a_3	a/g		Verified	genomic	614		ccaagaaaaggNtgatgagggttaa
479	IGR2181a_4	a/c		Verified	genomic	83		gtggaggctgaNagtaggcgagttt
480	IGR2182a_1	a/g		Verified	genomic	115		tgccccaagaaaaggNtgatgagggttaaacc
481	IGR2184a_1	other/w++	Poly t	Verified	genomic	58		tccttcatttttagccgaaagactcccttagcaNtttt
482	IGR2184a_4	a/t		Not yet verified	genomic	448		tgccatgttggNtgctgcaccc
483	IGR2184a_5	ins/del	ins/del t	Verified	genomic	380		tatttttttttaaglacNttaagttctagggt
484	IGR2185a_2	t/c		Not yet verified	genomic	420		gtctagatccNtgaggaaac
485	IGR2185a_3	t/g		Not yet verified	genomic	453		ttccacaatggNgaactagttt
486	IGR2186a_1	a/t		Not yet verified	genomic	184		gttcataacttNtccccgttt

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
487	IGR2188a_1	t/c		Not yet verified	genomic	549		tttgctgaagttgNttatcaacttaa
488	IGR2189a_1	a/g		Verified	genomic	475		atatgatgcattacNtttatcgatttg
489	IGR2189a_2	t/c		Verified	genomic	252		cctgtctgtgcNggtttcaa
490	IGR2190a_2	t/c		Not yet verified	genomic	343		ttattgccNcaatttc
491	IGR2190a_4	a/g		Not yet verified	genomic	326		ttgggtgataNgctattaatta
492	IGR2191a_1	a/g		Not yet verified	genomic	286		tgttgattttNgaigticc
493	IGR2191a_2	t/c		Not yet verified	genomic	353		actgcttgaatgNgcccagattc
494	IGR2191a_3	a/g		Not yet verified	genomic	390		ttgtctcttgtctcNttggtttcaaaa
495	IGR2191a_4	a/t		Not yet verified	genomic	498		gcggttttgaNtgagtttctt
496	IGR2192a_1	t/c		Not yet verified	genomic	515		tttttttgNtttcatttgc
497	IGR2192a_2	t/c		Verified	genomic	506		cccttcNttttttg
498	IGR2192a_3	t/g		Verified	genomic	359		tttatgaatcgggNgctctgtatt
499	IGR2193a_1	t/g		Verified	genomic	361		ttcaggagccttNtaaggcagg
500	IGR2193a_2	t/g		Verified	genomic	376		ggccctggNggtgacaaaa
501	IGR2193a_3	t/c		Verified	genomic	423		attttatttcNccttcactat
502	IGR2194a_1	a/g		Verified	genomic	57		cagagagatccNctgttagtctga
503	IGR2194a_2	a/g		Verified	genomic	196		agagatcttNgggtgtctc
504	IGR2194a_3	t/c		Verified	genomic	220		atttcctgaaNttgaatgttgccc
505	IGR2197a_1	other/w+		Not yet verified	genomic	498		gtctaactagtcccaNcgagatgagccgggt
506	IGR2198a_1	g/c		Verified	genomic	233		cagtagacgaacNalgcataatacca
507	IGR2198a_3	a/t		Not yet verified	genomic	98		tcctggggctttNacgttttagtg
508	IGR2199a_1	t/c		Verified	genomic	357		cagagataagaaNtagtttccaagaa
509	IGR2200a_1	a/c		Verified	genomic	176		acaggcttNgacagaggacttga
510	IGR2202a_1	t/c		Verified	genomic	308		tcactaaattctagaaaaNaaagattctaggcagt

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
511	IGR2202a_2	a/t		Not yet verified	genomic	330		taggcagttgctgNtattaaaaatcat
512	IGR2202a_3	a/g		Verified	genomic	528		caggactaaagtgaNctactctgaaaga
513	IGR2202a_4	ins/del	12 bp deletion	Not yet verified	genomic	622		tttttgacacacacaatgacactNcacttagagaagtc
514	IGR2203a_1	a/g		Verified	genomic	329		acaaacaaataaacaNiaaaacaaacccaca
515	IGR2203a_2	other/w+	Poly a	Verified	genomic	216		cagagtgattcigtgttNaaaaaaaaa
516	IGR2204a_1	ins/del		Not yet verified	genomic	584		acagcaaaaggccttNactgaaggactc
517	IGR2206b_1	t/g		Not yet verified	genomic	520		agggcggtgcagNagaagagctgggcc
518	IGR2207a_1	a/c		Verified	genomic	428		ggfataataattttNcgtcatcagaccic
519	IGR2209a_1	t/g		Verified	genomic	153		tgtgggggaaggNctatagccaagat
520	IGR2209a_2	g/c		Verified	genomic	234		gcacttctctcaaNctggagaccaccag
521	IGR2209a_3	a/g		Verified	genomic	462		ggccatcagaatctcNagttgatctttaa
522	IGR2210a_1	g/c		Verified	genomic	297		tcctgtaaggNtctgtagggcc
523	IGR2210a_2	t/c	c on ref. sequence	Verified	genomic	610		catctagggtgtaNgttccatgaggg
524	IGR2213a_1	g/c		Verified	genomic	314		cggctactgtggagcaNagagggtgctccaa
525	IGR2214a_1	a/t	a on ref. sequence	Verified	genomic	318		taaccaccaggctccagaNgtgcctagatccag
526	IGR2215a_1	a/g		Verified	genomic	198		agatciggagagattccccacNagagtcctatttccc
527	IGR2221a_1	other/w+		Not yet verified	genomic	214		cagagacttgtctgagNaaaaaaagaaaaa
528	IGR2221a_2	a/t		Verified	genomic	261		gaaaaaaaggaaaaaNatagcatgtta
529	IGR2221a_3	t/g		Verified	genomic	289		gctatcaataatcaaggccactgagNgtctatggatat
530	IGR2221a_4	ins/del		Not yet verified	genomic	231		aaaaagaaaaaNaagaaaaa
531	IGR2221a_5	t/c		Not yet verified	genomic	79		aaaaatlagccaagtgNgtggcaggcac
532	IGR2222a_1	a/g	g in ref. sequence	Verified	genomic	446		gcacatggggcacaNggcacactcacca
533	IGR2222a_2	t/c	c on ref. sequence	Verified	genomic	476		cagagtgcacgcgaNagcaccgccgcat

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
534	IGR2223a_1	ins/del		Not yet verified	genomic	194		ttttggttcctctctatttaaNaatggtatctttgtga
535	IGR2223a_2	other/w+	at repeat	Not yet verified	genomic	485		gcctcaaggNaagaatatt
536	IGR2224a_1	a/g		Verified	genomic	300		ctccaaccatgccNccctcttcctggggc
537	IGR2224a_2	t/c		Verified	genomic	387		gagtcctagtaaatggacNaccaagtaactaagac
538	IGR2224a_3	t/c		Verified	genomic	389		cctagtaaatggactaNaagtaactaagaccaa
539	IGR2224a_4	t/c		Verified	genomic	582		tgaggacatcacagNgtctccagaaaggla
540	IGR2225a_1	a/c		Verified	genomic	464		agtcgggtcicaNagtgcctcatgtatt
541	IGR2226a_1	a/g	g on ref. sequence	Verified	genomic	204		taaagagaaagaaNcattgtctctgatt
542	IGR2226a_2	t/g	t on ref. sequence	Verified	genomic	426		catgtctctatgtgtctNgccaaaggagctgaa
543	IGR2226a_3	t/c		Verified	genomic	524		ggaatgtctgtaaNgcatcatcagtg
544	IGR2226a_4	t/c		Verified	genomic	272		taaggagtagtatacNgtacaaaaglatct
545	IGR2228a_1	g/c	g on ref. sequence	Verified	genomic	450		galattcacagtagtgNggaagaccaacatta
546	IGR2229a_1	t/g	t on ref. sequence	Verified	genomic	298		ttttctgtgtgttNttttttccatcac
547	IGR2230a_1	t/c		Verified	genomic	608		catacttttagccaNttagggtgtatt
548	IGR2233a_1	g/c	g on ref. sequence	Verified	genomic	597		tgtgaaaccitgggNaagtatttaa
549	IGR2234a_1	a/g		Verified	genomic	362		taatcccagcaactcNgaggcgtgagaca
550	IGR2234a_2	g/c	g on ref. sequence	Verified	genomic	395		gaatccttgaaacctgNgaggcagaggttga
551	IGR2235a_1	t/c		Verified	genomic	153		gtgtctcacatgtNcatgtggccaagga
552	IGR2235a_2	t/g		Verified	genomic	386		agthaaagccttaNaattatacaaat
553	IGR2236a_1	a/g		Verified	genomic	256		ttacctagtaaacccgNtcacagatacatca
554	IGR2236a_2	ins/del		Not yet verified	genomic	321		atttgaattacggagtcagatNttggctctctact
555	IGR2236a_3	t/c		Verified	genomic	441		gaagggccaggcacaNgcttctctcagtcg
556	IGR2237a_1	a/g		Verified	genomic	395		agcaaggcctctaacNctfctctctaaaaatc
557	IGR2237a_3	a/g		Verified	genomic	619		tgggccaatgacccccNggctctttttgtgac
558	IGR2238a_1	a/g		Verified	genomic	92		ccctgctctctcNgggtcccaccctg
559	IGR2238a_2	a/c		Verified	genomic	115		accctggccaatgaNccccgggtctcttt

Table 3

SEQ. ID NO.	Polymorphism Name	Polymorphism type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
560	IGR2238a_3	ins/del		Not yet verified	genomic	247		gtccccactctactattNactcttccaacct
561	IGR2238a_4	a/g		Not yet verified	genomic	442		tgatctggctNcgccctgcctaaaca
562	IGR2239a_1	a/c		Not yet verified	genomic	256		ctgctctccgcactgNtgggcagtggtgg
563	IGR2240a_1	t/c		Verified	genomic	545		agtgcctattttgagaNagggcccgagcat
564	IGR2242a_1	t/g		Not yet verified	genomic	119		gtgggttaagattNgggtcagagctcta
565	IGR2243a_1	a/c	c in ref. sequence	Verified	genomic	256		tgccccctgtatNgaagagagggc
566	IGR2244a_1	other/w+	Poly t	Verified	genomic	220		ttttttttNggctccctgaccc
567	IGR2244a_2	g/c	c on ref. sequence	Not yet verified	genomic	73		ccaccagcctggNtaattttgt
568	IGR2244a_3	t/c	t on ref. sequence	Verified	genomic	469		gaggttaagNtccagggtctct
569	IGR2244a_4	t/c		Verified	genomic	576		tgagggtctctNcatcttctaaga
570	IGR2245a_2	a/g		Verified	genomic	145		aggacaaatgggNagggagtgaggag
571	IGR2245a_3	a/g	g in ref. sequence	Verified	genomic	397		attacaggcaccNccaccagcaggg
572	IGR2245a_4	t/g		Verified	genomic	434		atttttagcggaNacgaggttcacca
573	IGR2245a_5	t/c		Verified	genomic	574		tgctgtccaNaggtcggacag
574	IGR2245a_6	ins/del	Poly t	Verified	genomic	261		ttttttttNgagacggag
575	IGR2246a_1	ins/del		Verified	genomic	629		ccaccagccccctgccaNtattattta
576	IGR2248a_1	t/g	t on ref. sequence	Verified	genomic	145		ctagatgcagtgNtcagcaggccag
577	IGR2249a_1	a/g	a in ref. sequence	Verified	genomic	289		aactgaaNgttccaatttct
578	IGR2250a_1	t/c		Verified	genomic	145		ggctcagcaccacaNccagcagggtt
579	IGR2250a_2	t/c	t on ref. sequence	Verified	genomic	253		ttctgtgtgtgcNtggggccttca
580	IGR2250a_3	t/g	t on ref. sequence	Verified	genomic	302		acaccctaggctcacNgagaggcctcc
581	IGR2250a_4	t/c	c on ref. sequence	Verified	genomic	389		tatcaatgagggtctaNtactgtgtacttac

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
582	IGR2251a_1	g/c	c on ref. sequence	Verified	genomic	269		taatccagcttggNaggcagaagcagg
583	IGR2251a_2	a/t		Verified	genomic	360		aaacacaaaaattNgctgggcgcgtg
584	IGR2251a_3	a/g	g in ref. sequence	Verified	genomic	392		cagctactcggagNctgaggcaggag
585	IGR2251a_4	g/c		Verified	genomic	436		aggcgaagattgcaNtgagccaagaacg
586	IGR2251a_5	other/w+	Poly a	Verified	genomic	526		tgacagaggaggagactcgtctctcctNaaaaaa
587	IGR2252a_1	a/g		Verified	genomic	323		cccaactagagtaaNtccctggacacacag
588	IGR2252a_2	t/c		Verified	genomic	405		tggccatcaggagNgggagggccagactg
589	IGR2253a_1	a/g	g in ref. sequence	Verified	genomic	246		ccggctccagcccNagcgcgcgagaa
590	IGR2254a_1	t/g			genomic	68		agcgcgcgcctggggctcNgggaacgcgg
591	IGR2255a_1	a/c		Not yet verified	genomic	477		ttctagtagccNtattaataaaatt
592	IGR2255a_2	a/g		Verified	genomic	217		gaggctgggagctNtgacttttcatt
593	IGR2255a_3	other/w+	Poly a	Verified	genomic	387		tcagagctaactggNaaaaaaa
594	IGR2256a_1	t/c		Verified	genomic	510		atcatagtcaccgcagNcctgaactcctaagctt
595	IGR2256a_2	other/w+	Poly a	Verified	genomic	344		ttctcaggatttgNaaaaa
596	IGR2256a_3	a/g		Verified	genomic	179		tgaataataactttaNtggtataatttaa
597	IGR2257a_1	a/g	g in ref. sequence	Verified	genomic	423		atataaigtgttgNgtaaagaatat
598	IGR2257a_2	t/c	t on ref. sequence	Verified	genomic	508		cagcagatttttaaNaaggaaatctaa
599	IGR2257a_3	g/c		Verified	genomic	621		ctattctacttcNgaagatggatgg
600	IGR2258a_1	other/w+	Poly t	Verified	genomic	575		tgcanttttttt
601	IGR2259a_1	other/w+	Poly t	Verified	genomic	234		gctaNttttttg
602	IGR2260a_1	t/c		Not yet verified	genomic	582		tcaacaataNgttaaatataa
603	IGR2261a_1	t/c		Verified	genomic	608		ggctgaggaggNggatcacc
604	IGR2262a_1	ins/del	Poly a	Verified	genomic	332		aagactccgtctcNaaaaaa
605	IGR2262a_3	a/g	a in ref. sequence	Verified	genomic	425		ttcagagcNtctgtccag
606	IGR2263b_1	g/c	g on ref. sequence	Verified	genomic	411		ttaagtgattctNcigtctcagcctcc

Table 3

SEQ. ID NO:	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
607	IGR2264a_1	other/w+		Verified	genomic	318		taatagctgttttttNtgcaaaatcactgt
608	IGR2265a_1	t/c	c on ref. sequence	Verified	genomic	249		cccacaattNggctcaa
609	IGR2265a_2	t/c	t on ref. sequence	Verified	genomic	340		gtagagaaaNgtaaatt
610	IGR2269a_1	a/g		Not yet verified	genomic	270		tatgtacaagtatctNtttgagctactgtct
611	IGR2272a_1	a/t		Verified	genomic	540		tttttaaaaaaaNttttaaggcatagga
612	IGR2272a_2	t/c		Verified	genomic	163		ctcttgaaggctgNggcaggaagatgc
613	IGR2272a_3	a/g	g in ref. sequence	Verified	genomic	384		tacaaaaatacaaaaaaattagccNggcgtgtg
614	IGR2272a_4	t/c		Verified	genomic	395		ttagccgggctgtgtgNgggcacctgtagtaccc
615	IGR2272a_5	g/c		Not yet verified	genomic	462		tttgaaccccgaggcggaNgttgaatgagtgagatt
616	IGR2273a_1	other/w+	Poly t	Verified	genomic	388		cccttatccacagNttttttttt
617	IGR2274a_1	t/g		Verified	genomic	311		tctccatgcaccgcaNtcacattgtgtgtg
618	IGR2274a_2	g/c		Verified	genomic	381		tcattagcctggcttNcattctcttctgaac
619	IGR2274a_3	t/c	c on ref. sequence	Verified	genomic	539		atactactatggNcctttgtctccg
620	IGR2276a_1	t/c	c on ref. sequence	Verified	genomic	113		cactactcatcttcNtgagcacaaaag
621	IGR2276a_2	a/c	c in ref. sequence	Verified	genomic	359		aaatgagtagccttcNtttgagagacagag
622	IGR2277a_1	a/g	a in ref. sequence	Verified	genomic	143		gatcatcicaagggttcNcaaaatcaagct
623	IGR2277a_2	other/w+	Poly t	Verified	genomic	485		gaigcaagaaNttttttttttt
624	IGR2279a_1	a/g	a in ref. sequence	Verified	genomic	165		acaggcatccaccaccNtgccctggtaattt
625	IGR2279a_2	t/c	c on ref. sequence	Verified	genomic	256		catgtgatctgccNgcctcagccttccaaa
626	IGR2279a_3	other/w+	Poly t	Verified	genomic	310		ccaatgcgcctggccNttttt
627	IGR2279a_4	g/c		Verified	genomic	108		cctctgcctccagggttNaagcagttctcctg
628	IGR2279a_5	t/c		Verified	genomic	277		gccttccaaagtgcNaggattacaggt
629	IGR2281a_1	a/c		Verified	genomic	144		catcttgcattatataaagaataac

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
630	IGR2281a_2	other/w+	Poly t	Verified	genomic	87		aaattaatttttctccNitttttt
631	IGR2281a_3	a/t		Not yet verified	genomic	72		taattttttaaatNaattttttcttc
632	IGR2283a_2	ins/del	ins/del 2bp	Verified	genomic	574		cctggctctcNttagttatt
633	IGR2284a_1	t/c	t on ref. sequence	Verified	genomic	486		gccttcactttccaNatcaccaatcagc
634	IGR2284a_2	t/c		Not yet verified	genomic	536		tgccaaglaactattNtaactctgagaatac
635	IGR2285a_1	a/c		Verified	genomic	171		gaaaaatgaagcNggagaaaaatgaa
636	IGR2286a_1	a/c		Not yet verified	genomic	261		tgctacatgcNagacaatca
637	IGR2287a_1	t/c	t on ref. sequence	Verified	genomic	107		cttggaggcgNgaggcaggcaga
638	IGR2287a_2	other/w+	Poly a	Verified	genomic	188		glgaaacccccgttctctactataaaaatacNaaaaaaaaa
639	IGR2287a_3	other/w+	Poly a	Verified	genomic	351		acagagcgagactccgctctcNaaaaaaaaa
640	IGR2287a_4	t/c		Verified	genomic	463		tgtgaaggactgggNtttcaaaaaatctg
641	IGR2287a_7	t/c		Verified	genomic	463		tatagaccattgNaaggactggg
642	IGR2288a_1	t/c	t on ref. sequence	Verified	genomic	170		atggcaaaaagaNtttattgaca
643	IGR2288a_2	a/g	a in ref. sequence	Verified	genomic	198		ggatgtggagtacNagaggaagagcagcc
644	IGR2289a_1	a/g	g in ref. sequence	Verified	genomic	215		cccaagtagctgggactNcagggtgtgccacca
645	IGR2291a_1	g/c	c on ref. sequence	Verified	genomic	153		cigtatcctagctacttNggaggctgaggcatga
646	IGR2291a_2	a/g		Not yet verified	genomic	548		tagcaagaagtNggaggagggtt
647	IGR2292a_1	a/c		Verified	genomic	256		gtctcatgtNatccccacc
648	IGR2292a_2	t/c	c on ref. sequence	Verified	genomic	589		tctatttatcttNaatttcctatt
649	IGR2292a_3	a/t		Verified	genomic	282		atggaattgtatcNtccctctttacaga
650	IGR2293a_3	ins/del	gt repeat	Verified	genomic	437		tgtgttgtNgtgtgtgtgtgtg
651	IGR2294a_1	t/c	c on ref. sequence	Verified	genomic	390		ccitggaaaaaNgggacactcc
652	IGR2294a_2	t/c		Verified	genomic	440		ttagcaaatggNacaccagga

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
653	IGR2294a_3	t/g		Verified	genomic	481		tgcacagatccNatgtccatgga
654	IGR2295a_1	ins/del		Not yet verified	genomic	438		attgtctgtcNgcaatatttgc
655	IGR2295a_2	a/g		Verified	genomic	535		tgcagctgagggNccicacgtgtagaa
656	IGR2297a_1	g/c	g on ref. sequence	Verified	genomic	65		taacicaagaaNattagagaaa
657	IGR2297a_2	t/c	c on ref. sequence	Verified	genomic	198		aaaacacicNtcaggata
658	IGR2297a_3	t/g	g on ref. sequence	Verified	genomic	487		ttctaaagaaaaNaattttcaaccca
659	IGR2297a_4	t/c	c on ref. sequence	Verified	genomic	588		gatttgcaccacNaggcctgccctaaaaga
660	IGR2297a_5	a/c	c in ref. sequence	Verified	genomic	446		ccctacaagccNgaagagag
661	IGR2298a_1	a/g	g in ref. sequence	Not yet verified	genomic	293		tttaaatgtaaatggNctaaatgctcca
662	IGR2299a_1	a/g	g in ref. sequence	Not yet verified	genomic	592		caaagacacaacNtgcagaatct
663	IGR2300a_2	t/c		Verified	genomic	606		ccaataacaggNtctgaaattg
664	IGR2303a_1	t/c	c on ref. sequence	Verified	genomic	189		ttttgtatctacNggcaaaatata
665	IGR2303a_2	g/c	g on ref. sequence	Verified	genomic	495		aatatctcattagtNataatgagccc
666	IGR2304a_1	t/c		Verified	genomic	483		cttggatgttNgaatggcat
667	IGR2304a_2	a/g		Verified	genomic	667		ggttgagtgtagacaNtacagggtaaaaa
668	IGR2305a_1	a/t		Verified	genomic	253		tttctgtaggaatNctgcataatcatttgg
669	IGR2308a_1	t/c	c on ref. sequence	Verified	genomic	339		tttgtatcctttglaagaaacNgctaglggcca
670	IGR2308a_2	a/g		Verified	genomic	561		taggtattgtcaaaaattgNactgcattataggaca
671	IGR2309a_2	t/c		Verified	genomic	610		gatgtgttttttNtggagacgg
672	IGR2310a_1	ins/del		Not yet verified	genomic	273		aaattttgtatttttNtagtagagatgggt
673	IGR2311a_1	a/g	g in ref. sequence	Verified	genomic	181		gcccagctcggagatgcNgtggcatgatgttgg

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
674	IGR2311a_2	a/c	c in ref. sequence	Verified	genomic	207		ttggctacigcaaNctccaccctccggg
675	IGR2311a_3	ins/del		Not yet verified	genomic	525		caaccctcgcctcctgggtNgcagttctccigcct
676	IGR2313a_1	other/w+	poly A	Not yet verified	genomic	499		aaaaaaaaNcaactaag
677	IGR2313a_2	a/g		Not yet verified	genomic	370		taaccaggNtgttcagg
678	IGR2313a_3	a/g		Not yet verified	genomic	335		aaatgggggNtgggaggaca
679	IGR2313a_4	t/c		Not yet verified	genomic	531		cagattaaNcagtaaat
680	IGR2313a_5	ins/w+	C insert	Not yet verified	genomic	391		agtttggcNatgatagg
681	IGR2314a_1	ins/w+	varying number of GT repeats	Verified	genomic	560		atgtttcaNgtgtgtgt
682	IGR2315a_1	a/g		Not yet verified	genomic	369		cttcattgcNaagagttgc
683	IGR2315a_2	t/c		Not yet verified	genomic	533		taatttctaNgcctgtcttt
684	IGR2315a_3	a/g	G in ref	Not yet verified	genomic	211		aacatgccNctgaaaca
685	IGR2316a_1	a/g	G in ref	Not yet verified	genomic	499		cccaggcttNtaggatga
686	IGR2316a_2	t/c	G in ref	Not yet verified	genomic	565		aaacccctgNtccgtataa
687	IGR2316a_3	a/g		Not yet verified	genomic	469		tgaataaNccccagtc
688	IGR2321a_1	t/c		Not yet verified	genomic			ttgtgaaaaNgtcaaatag
689	IGR3000a_1	a/g	a in ref. sequence	Verified	genomic	224		ttttagaaNtgatacttt
690	IGR3000a_2	a/g		Verified	genomic	558		ttaagaaatatgtNtttcttactatc

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	Comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
691	IGR3002a_1	g/c	g on ref. sequence	Verified	genomic	146		ctggcagNggtcga
692	IGR3002a_2	a/g		Verified	genomic	511		atattgaacNacatagat
693	IGR3003a_1	a/g		Verified	genomic	494		tgaacccccNtcttactt
694	IGR3004a_2	t/c	c on ref. sequence	Verified	genomic	430		gagtggaactctcacNgcccagatttctc
695	IGR3004a_3	ins/del		Verified	genomic	285		atttctctcttNtttcttctt
696	IGR3005a_1	t/c	t on ref. sequence	Verified	genomic	494		aggagtagNttagatagaa
697	IGR3005a_2	g/c	c on ref. sequence	Verified	genomic	538		agtagcacNactaccca
698	IGR3005a_3	a/c	c in ref. sequence	Verified	genomic	34		ccatgaaggcaccacaaNtcaactgccaggt
699	IGR3006a_1	other	gt repeat	Verified	genomic	234		ccagttctgacgatacNtctgtgtgtg
700	IGR3006a_2	t/g		Verified	genomic	227		cagttctgacNatacgt
701	IGR3007a_1	t/c		Verified	genomic	458		cgtgaagccaNtgcgcca
702	IGR3007a_2	other/w+	Poly t	Verified	genomic	176		aaatactgtaccctgtgacNtttt
703	IGR3008a_1	t/c		Verified	genomic	147		cacttattaNttaccata
704	IGR3008a_2	a/c		Verified	genomic	339		tgcatagcaaNtctcactt
705	IGR3008a_3	t/c	C in ref. sequence	Verified	genomic	342		atgcaactcNcacttcacc
706	IGR3013a_1	other	gt repeat	Verified	genomic	637		cacattatataatgcNtgtgtgtg
707	IGR2016a_1	a/g	a in ref. Sequence	Verified	genomic	636		ctgtgtgtacagcNtgtgttcattttgc
708	IGR3018a_1	a/g	G in ref. sequence	Verified	genomic	238		gggcactgacaccccNcctgtgtggggccc
709	IGR3018a_2	t/g		Verified	genomic	191		gggcacctgtgtcNtgcattgttcttta
710	IGR3019a_1	a/g		Verified	genomic	205		tgtgttagaataatttgcNtattgtaggctaatgta
711	IGR3019a_2	a/g		Verified	genomic	388		cagctttattgaagaNgcaatgttacag
712	IGR3020a_1	g/c	g on ref. sequence	Verified	genomic	172		gtctctgccctggctNtgttttagctgtgcc
713	IGR3020a_2	a/c	c in ref. sequence	Verified	genomic	349		cttacttagccctagaNaacaaattataag
714	IGR3020a_3	a/g	a in ref. sequence	Verified	genomic	542		tataggaacttacNataatgttaggtca

Table 3

SEQ. ID NO:	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
715	IGR3022a_1	a/t	t on ref. sequence	Verified	genomic	267		gctggagagctgNctcatactgagcag
716	IGR3023a_1	g/c	c on ref. sequence	Verified	genomic	79		tctcttagggcaNagtgcaggctccc
717	IGR3023a_2	ins/del	ct repeat	Verified	genomic	264		attctctctctNtctctctgatag
718	IGR3023a_3	t/g	g on ref. sequence	Verified	genomic	368		ggcatgatcatatagcNcactgtaacttg
719	IGR3023a_4	t/c	t on ref. sequence	Verified	genomic	580		gggattacagggtgtgaaNcaccatacctggctaa
720	IGR3029a_1	a/g	A in ref. sequence	Verified	genomic	201		ctggagggtNcacagacagg
721	IGR3029a_2	a/g	A in ref. sequence	Verified	genomic	498		agtcacagNacaaagct
722	IGR3030a_1	t/c	T in ref. sequence	Verified	genomic	181		attcatgcNtgcctttt
723	IGR3032a_1	a/g	A in ref. sequence	Verified	genomic	158		actagggaNgccaaggc
724	IGR3032a_2	g/c	C in ref. sequence	Verified	genomic	281		glaatcccaNctattcggg
725	IGR3035a_2	a/g	G in ref. sequence	Verified	genomic	223		gtcagggggaNtgatggaaa
726	IGR3036a_1	a/g		Verified	genomic	187		aaatacaNaaaataa
727	IGR3037a_1	a/c		Verified	genomic	521		gggagaaccNiccacag
728	IGR3038a_2	t/c		Not yet verified	genomic	464		aaatacagaaaNactttttgtgtt
729	IGR3039a_1	a/g	G in ref. sequence	Verified	genomic	387		gggccagaggNtggaagcgaag
730	IGR3040a_2	t/c		Verified	genomic	517		cccgtcacaNaggggagg
731	IGR3040a_3	t/c		Verified	genomic	133		acctgagaaNccaacacaacga
732	IGR3041a_1	t/c	T in ref. sequence	Verified	genomic	331		tcgtcacaNgggaagat
733	IGR3042a_1	a/g		Verified	genomic	435		caccctagaNatgatggaa
734	IGR3043a_1	t/c		Verified	genomic	278		ataggcccagtgatggNggcctggcacatgaact
735	IGR3044a_1	t/c		Verified	genomic	477		caggcatcaatgcagaNttagtgtttttcaggg
736	IGR3044a_2	t/c		Verified	genomic	513		ctctggcagactttttcNctgtcacatcctccca

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
737	IGR3045a_1	t/g		Verified	genomic	137		aagcatggagcagtgtaacNcaaggacctgtggaata
738	IGR3046a_1	a/g		Verified	genomic	293		tgtggccacagtgccctNgccagggtccaagcc
739	IGR3047a_1	t/g		Verified	genomic	455		cagactctctccctNggccaggatattgccttgt
740	IGR3047a_2	a/g		Verified	genomic	522		ttagctggccctgcccNggactggggagagtaa
741	IGR3047a_3	t/c		Verified	genomic	609		gtagctctctactcNgcctgcattacatagca
742	IGR3049a_1	t/c		Verified	genomic	437		tttatcacacctNatttcagcagacacaga
743	IGR3049a_2	t/c		Verified	genomic	611		gtccacggccctgcccNttgccagacgggctcca
744	IGR3050a_1	t/c		Verified	genomic	224		ttcgaatactgagatcNgaagaagtgctcc
745	IGR3051a_1	a/t		Verified	genomic	667		tttagagatagaaggaaNggagggttagat
746	IGR3053a_1	a/g		Verified	genomic	364		ggggctcttagaaaNggctttcttagaa
747	IGR3053a_2	t/c		Verified	genomic	481		gttaacagctgacatggNggccacagtgaggagaca
748	IGR3054a_1	other/w+	poly A	Verified	genomic	597		cctagtgaaatggNaaaaaaaaa
749	IGR3055a_1	t/c		Verified	genomic	375		ccctctccaccatNctccagcagaaaggacag
750	IGR3055a_2	other/w+	Poly a	Verified	genomic	133		aaaaaaaaaaaaaNttgcttaacatt
751	IGR3056a_1	a/g		Verified	genomic	328		cttcaaaaaagatgacaNlaataacctgctctagg
752	IGR3056a_2	a/g		Verified	genomic	383		aaatacagtgagcNctgacacattacaggcc
753	IGR3057a_1	t/g		Verified	genomic	549		ttagcagtcactcctcattcNctactcctctagccctg
754	IGR3059a_1	a/t		Not yet verified	genomic	94		tatatatatatatNlattcaagggttggtgctata
755	IGR3059a_2	other/w+	at repeat	Verified	genomic	63		caacaacNtatatatatatata
756	IGR3060a_1	g/c		Verified	genomic	102		tccactgttaaggNctcttggaattcttt
757	IGR3061a_1	t/c		Verified	genomic	362		tttcaattatgtatataNttttactccagaagt
758	IGR3061a_2	t/c		Verified	genomic	592		caatatgtcatcaNacttttaaaagcatgacttc
759	IGR3062a_2	t/c		Verified	genomic	139		tgaacatatattataaNggctgcctatgccitaa
760	IGR3064a_1	t/c		Verified	genomic	616		cttgccaaggtatagNgcactttcttgaataaa
761	IGR3065a_1	a/g		Verified	genomic	358		tttatccatttttaatacaNgtgctttttattgctgag
762	IGR3066a_1	a/t		Verified	genomic	351		tctggaagtgcgcctcNaccigccctccagctctg
763	IGR3068a_1	t/c		Not yet verified	genomic	337		gaagttcccNgttagcagggg
764	IGR3072a_1	ins/del	caaa repeat	Verified	genomic	383		caacaacaaacaaacaaacaaNaactagccgggcatg
765	IGR3072a_2	a/t	t on ref. sequence	Verified	genomic	578		taaaataaaataaaaaNaaaacgaaaaataattt
766	IGR3078a_1	a/g		Verified	genomic	313		gggcaggaggtggNcaagcactagag
767	IGR3079a_1	a/g		Not yet verified	genomic	120		ccctcgaataaaagtcNctctcagtatatac

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference sequence	Gene Name	Flanking Sequence
768	IGR3081a_1	t/g	t on ref. sequence	Verified	genomic	317		gagtcctattcttctNggggtgcacaccg
769	IGR3081a_2	a/g		Verified	genomic	286		gaaacgaccagNaatgcgcctcgcg
770	IGR3083a_1	g/c	g on ref. sequence	Not yet verified	genomic	397		gctcgggcccgtNgcccgggcccagacccca
771	IGR3084a_1	t/c	t on ref. sequence	Verified	genomic	504		cggcaggcgtNcagagcttt
772	IGR3086a_1	a/t		Verified	genomic	234		ttgagatggttNtggcgatgacc
773	IGR3087a_1	other/w+	Poly t	Verified	genomic	325		ggaacaatctcNtttt
774	IGR3087a_2	ins/del		Verified	genomic	90		ttccagattNgacataa
775	IGR3087a_3	t/c		Verified	genomic	185		gtaataaaNctctatcg
776	IGR3088a_1	other/w+	Poly t	Verified	genomic	108		tgataagtcctgcNtttttt
777	IGR3088a_2	a/t	t on ref. sequence	Verified	genomic	269		gcaaacaccNccacacca
778	IGR3089a_1	a/g		Verified	genomic	559		ctagaacacaaaaNgtagaacaaaa
779	IGR3090a_1	g/c		Verified	genomic	558		agttgctaNaacatctgt
780	IGR3095a_1	other/w+	Poly a	Verified	genomic	257		actcgtctcNaaaaacaaaaa
781	IGR3095a_2	a/t		Not yet verified	genomic	178		aaattgcttNacccggaggc
782	IGR3096a_1	t/c	t on ref. sequence	Verified	genomic	316		ccTggagaaNagctgagaa
783	IGR3096a_2	a/g		Verified	genomic	406		aggTggcacNgatctctaaa
784	IGR3096a_3	t/c		Verified	genomic	424		aaagctgtccNgcTgcca
785	IGR3097a_1	a/g		Verified	genomic	338		agaaatcatgagagcagNaaaggagaaagggt
786	IGR3097a_2	a/c		Verified	genomic	472		acaacaacacaaNaaaaagagcTcaaatgg
787	IGR3098a_1	a/t		Verified	genomic	373		gtctttgtaaaaacNacaaatttata
788	IGR3100a_1	t/c		Verified	genomic	243		ggcaggcggatcaNgaggTcaagagatccaga
789	IGR3103a_1	t/c		Verified	genomic	326		aaagggcNgacatggc
790	IGR3105a_1	t/c		Verified	genomic	231		agtggTgNgatctgg
791	IGR3105a_2	a/g		Verified	genomic	575		ttaccatNtaacccaa
792	IGR3105a_3	ins/w+	CA repeat	Verified	genomic	187		TgtgTgNagacagaatcttg
793	IGR3108a_1	t/c		Verified	genomic	348		ggTcccNggccagg
794	IGR3110a_1	ins/w+	GAA repeat or deletion	Verified	genomic	59		ggaagaNgaagaag
795	IGR3111a_1	a/g		Verified	genomic	199		ctgaggNgTggTgcct

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
796	IGR3112a_1	t/c		Verified	genomic	72		ttactttgNccagcttcc
797	IGR3113a_1	g/c	G in ref	Not yet verified	genomic	321		aatggatNtatgtcaga
798	IGR3113a_2	a/t	T in ref	Not yet verified	genomic	368		agggacccNaatagttt
799	IGR3113a_3	t/g	T in ref	Not yet verified	genomic	477		attcagaNgigtgt
800	IGR3114a_1	a/g		Verified	genomic	571		cacaagtNtccacagag
801	IGR3115a_1	other/w++	Poly t	Verified	genomic	557		gaatgatgcNtttttttt
802	IGR3117a_1	t/c		Verified	genomic	452		cccaaatgNtacctat
803	IGR3118a_1	t/c	C in ref	Not yet verified	genomic	116		tggcataNagaaggt
804	IGR3119a_1	a/c		Verified	genomic	301		gcctagatcNctgtctgca
805	IGR3119a_2	a/c		Verified	genomic	534		ggccatggtNlatggcc
806	IGR3121a_1	g/c	G in ref	Not yet verified	genomic	586		agtactggNaccctgggc
807	IGR3122a_1	t/g	G in ref	Not yet verified	genomic	144		catgtcNactacact
808	IGR3122a_2	a/g		Verified	genomic	441		gtaccagcNgctagtgga
809	IGR3125a_1	del/w+	del of ACTGT from ref.	Verified	genomic	3121		aaatgggNactgtctcg
810	IGR3125a_2	a/c	C in ref	Not yet verified	genomic	384		ggcaaacNcaccacg
811	IGR3129a_1	t/c		Verified	genomic	193		ccgtggaNttgggt
812	IGR3131a_1	t/g		Verified	genomic	1308		tgtgtgNggggtctga
813	IGR3133a_1	a/g		Verified	genomic	2029		tcgggcaNgcattgca
814	IGR3133a_2	t/g		Verified	genomic	2301		ttttagtNtgagctcc
815	IGR3134a_1	a/g		Verified	genomic	2594		gcctcaNtggatttt
816	IGR3138a_1	a/g		Verified	genomic	508		agctctNccctcag
817	IGR3141a_1	t/c		Verified	genomic	125		gggaagcNggctggg
818	IGR3145a_1	a/c		Verified	genomic	373		gaaaggaNtgaatgc
819	IGR3145a_2	a/g		Verified	genomic	420		gcctgcaNcacttgc
820	IGR3145a_3	ins/w++	Polymorphic CAAA	Verified	genomic	379		aaaggactgaaaNgccccagagc

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
821	IGR3148a_1	t/c		Verified	genomic	1733		ttcacaccNggaagct
822	IGR3149a_1	a/g		Verified	genomic	2119		aggaaacNttctct
823	IGR3150a_1	a/g		Verified	genomic	262		ccagagaNgtacagaa
824	IGR3152a_1	t/c		Verified	genomic	1223		gccgggcNgatagcct
825	IGR3153a_1	other/w+	CA repeat	Verified	genomic	1817		ttgcgtaNacacaca
826	IGR3153a_2	g/c		Verified	genomic	2055		gaactggaNagaagtctc
827	IGR3158a_1	a/c		Verified	genomic	669		gggagacNccttttac
828	IGR3159a_1	ins/w+		Verified	genomic	1004		gtgtgtNgggggg
829	IGR3161a_1	g/c		Verified	genomic	2042		gctgagaNcctatcat
830	IGR3162a_1	a/g		Verified	genomic	2539		gcctctNtatgcag
831	IGR3162a_2	a/g		Verified	genomic	2686		tgggacaNgaacaac
832	IGR3162a_3	a/g		Verified	genomic	2816		cctcccNtgaggccc
833	IGR3162a_4	g/c		Verified	genomic	2959		cagccccNtctccc
834	IGR3163a_1	t/c		Verified	genomic	478		gacagtaNagcctgtga
835	IGR3166a_1	a/g		Verified	genomic	325		gctgctatNaggtgcagg
836	IGR3166a_2	other/w+	poly t	Verified	genomic	651		ccatccttcNtttttt
837	IGR3169a_1	a/g		Verified	genomic	40		tgtcgcccNagtcacagt
838	IGR3169a_2	t/c	t in ref	Not yet verified	genomic	505		aacggagtNgtgcctct
839	IGR3170a_1	t/g		Verified	genomic	73		gtgtggaNagttaaa
840	IGR3170a_2	t/c		Verified	genomic	191		cagtcccNgagaagt
841	IGR3171a_1	t/c		Verified	genomic	213		tctattaaNgggagaatcc
842	IGR3173a_1	a/g		Verified	genomic	126		taaatcccaNattgattc
843	IGR3174a_1	a/c		Verified	genomic	489		aataaataNtcaattatt
844	IGR3176a_1	other/w+	poly t	Verified	genomic	107		cccccaaccNtttttt
845	IGR3178a_1	a/g		Verified	genomic	310		ggatcatNtttaagag
846	IGR3178a_2	del/w+		Verified	genomic	279		ggaaattgtNaatactt
847	IGR3179a_1	t/c	C in ref	Not yet verified	genomic	554		tccfgggcNtcaagggga
848	IGR3181a_1	a/g	G in ref	Not yet verified	genomic	257		gggaccaNgcagaaa

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
849	IGR3182a_1	a/c	C in ref	Not yet verified	genomic	614		acaaacNcaacaa
850	IGR3182a_2	a/t		Verified	genomic	620		aacacaacNaaaaaacaggccaa
851	IGR3183a_1	t/g		Verified	genomic	131		aaacaggNccaaatgacta
852	IGR3183a_2	other/w+	poly A	Verified	genomic	248		agggaagNaaaaaa
853	IGR3185a_1	a/t		Verified	genomic	211		aaacaatcNctacagtt
854	IGR3185a_2	g/c	G in ref	Not yet verified	genomic	558		aaaacatNatacagga
855	IGR3185a_3	a/g	G in ref	Not yet verified	genomic	529		aaaagtNaagtccta
856	IGR3186a_1	other/w+	poly t	Verified	genomic	174		gttgcaaNttttttt
857	IGR3186a_2	g/c		Not yet verified	genomic	58		aaaacatNatacagg
858	IGR3188a_1	a/g		Verified	genomic	152		gcactaNccaaaatat
859	IGR3188a_2	a/g	A in ref	Not yet verified	genomic	508		aacaataNcaaatgtt
860	IGR3189a_1	a/g	G in ref	Not yet verified	genomic	139		acctaggNatttcactcc
861	IGR3189a_2	g/c		Verified	genomic	394		gaaagaaNataatgtt
862	IGR3189a_3	a/g	A in ref	Not yet verified	genomic	429		gttcaaNatctgac
863	IGR3191a_1	a/g		Verified	genomic	215		ccccgcNaccctacitt
864	IGR3191a_2	t/c		Verified	genomic	246		cacttagNctttatc
865	IGR3192a_1	t/c		Verified	genomic	328		gggagaNccacacct
866	IGR3193a_1	a/g		Verified	genomic	409		gagacaNgagagga
867	IGR3194a_1	t/c	C in ref	Not yet verified	genomic	296		aaaggatNtgggtctt
868	IGR3196a_1	t/c		Verified	genomic	137		cacacgtNgcgtatgca
869	IGR3196a_2	other/w+	tg repeat	Verified	genomic	177		gcataaaaNgigtgtgtgt
870	IGR3197a_1	other/w+	poly t	Verified	genomic	308		tgtgggaNttttttt

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphism details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
871	IGR3199a_1	other/w+	poly a	Verified	genomic	131		tgagctcNaaaaaa
872	IGR3200a_1	a/g		Verified	genomic	342		ctccaccNttgtccccc
873	IGR3201a_1	a/t	A in ref	Not yet verified	genomic	234		aggactNattctcta
874	IGR3203a_1	a/t		Verified	genomic	538		ctgtctcNaggagcca
875	IGR3205a_1	t/g		Verified	genomic	1035		taatggaNlaaggat
876	IGR3205a_2	a/g		Not yet verified	genomic	548		cacatggttNcaatgtcac
877	IGR3206a_1	t/g		Verified	genomic -	1176		aagatctcNaggggtggg
878	IGR3206a_2	g/c		Verified	genomic	1511		gacaggNattgtctat
879	IGR3206a_3	g/c		Verified	genomic	552		gagacaggNattgtctat
880	IGR3207a_1	t/c		Verified	genomic	205		aaaaaaccttNggctgtct
881	IGR3208a_1	a/t		Not yet verified	genomic	206		ctgcctcgttNttacctggg
882	IGR3210a_1	other/w+		Verified	genomic	59		ttttgtgtgtgtgtgtgtgtgt
883	IGR3222a_1	a/c		Not yet verified	genomic	594		gccacatNtgcataca
884	IGR3230a_1	t/g		Verified	genomic	462		agagacacaccctgggNagagatgctgg
885	IGR3230a_2	a/g		Verified	genomic	491		cccacitccaaccNtgcctgg
886	IGR3236a_1	t/g		Not yet verified	genomic	320		gagaggNtgaagt
887	IGR3238a_1	g/c	G in ref	Not yet verified	genomic	280		ggggccagNgcaagtt
888	IGR3242a_1	a/g		Not yet verified	genomic	319		aacctaNggggagggg
889	IGR3244a_1	ins/w+		Not yet verified	genomic	91		tcccgctNtgtgtct
890	IGR3248a_1	t/c		Not yet verified	genomic	395		actgtcNccaactt
891	IGR3252a_1	other/w+	poly T	Not yet verified	genomic	183		tctaccNttttttt
892	IGR3252a_2	t/g		Verified	genomic	136		glgctgtNggacigaa

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
893	IGR3266a_1	a/g		Not yet verified	genomic	156		caggTggNatgtcttg
894	IGR3266a_2	a/g		Verified	genomic	185		ccactgggNccctggcct
895	IGR3268a_1	t/c		Not yet verified	genomic	367		tgtttgtaNtctctcc
896	IGR3268a_2	t/c		Not yet verified	genomic	385		aggaacTgNctcgacat
897	IGR3274a_1	a/g		Verified	genomic	126		ctctaggaNcatttcag
898	IGR3274a_2	a/g		Verified	genomic	284		tgagaaatNcagtgagc
899	IGR3276a_1	other/w+	poly t	Verified	genomic	340		ctctctctcNttttttttt
900	IGR3276a_2	ins/w+	ins G	Verified	genomic	221		ttctctgcctNtgatttgg
901	IGR3278a_1	other/w+	poly t	Verified	genomic	69		ctctctctcNttttttttt
902	IGR3292a_1	other/w+	poly t	Verified	genomic	518		ttcatctcNttttttttt
903	IGR3294a_1	t/c	T in ref	Verified	genomic	366		aatatctcctcttcaagaaNttgaattttgaatct
904	IGR3298a_1	a/g		Not yet verified	genomic	76		gttccttcogNttttccac
905	IGR3300a_2	g/c		Verified	genomic	366		cctgagaggcatcaaaaNtcaaatatgatcaa
906	IGR3302a_1	a/g		Verified	genomic	218		catctattgtatcNgtcacctgattggg
907	IGR3304a_1	ins/w+		Verified	genomic	319		tttggagggtgaggTgggNggatcaggaggTcaggag
908	IGR3310a_1	a/g		Not yet verified	genomic	361		ggaccaaNctggggTg
909	IGR3310a_4	other/w+	poly a	Not yet verified	genomic	86		aaaaaaaaaNgtttcccc
910	IGR3312a_1	ins/w+		Verified	genomic	249		cagataagcatcagatttgNaaactacaatgggaatg
911	IGR3324a_1	t/c		Not yet verified	genomic	388		tattgtattccaaatNtggatgtagccacca
912	IGR3326a_1	other/w+	tg repeat	Verified	genomic	261		tattctagtcTgagaggNttttgtttgtttg
913	IGR3326a_3	t/g		Verified	genomic	295		ttgtttNtttagacagagtcctca
914	IGR3328a_1	t/c		Verified	genomic	586		tatatattgttttcatNgtgaattctctttgacc
915	IGR3330a_1	other/w+	poly t	Verified	genomic	200		tacctcNtttttt

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
916	IGR3330a_2	other/w+	poly t	Verified	genomic	512		tttttttNaaccttaaa
917	IGR3332a_1	other/w+	poly a	Verified	genomic	347		catctcNaaaaaa
918	IGR3336a_2	other/w+	poly a	Verified	genomic	467		aaaaaaaaNgagagag
919	IGR3340a_1	a/g		Not yet verified	genomic	524		agactaNcacagaaa
920	IGR3348a_1	t/c	T in reference	Verified	genomic	91		ccacatNctctcc
921	IGR3348a_2	a/g	A in reference	Verified	genomic	202		gatgggNagcatt
922	IGR3348a_3	a/g	A in reference	Verified	genomic	216		tttcatNtgtttt
923	IGR3348a_5	t/c		Verified	genomic	129		aatgatNgccatt
924	IGR3348a_6	a/g		Verified	genomic	264		gttcatNtctctg
925	IGR3348a_8	a/g	A in ref	Verified	genomic	618		gatttttNataagg
926	IGR3348a_9	a/g		Not yet verified	genomic	584		tctaacNtttaag
927	IGR3350a_1	g/c	G in Reference	Verified	genomic	77		catgagNatggaa
928	IGR3350a_10	a/g		Not yet verified	genomic	451		ccTgcctNatggccc
929	IGR3350a_11	t/c	C in reference	verified	genomic	450		ccTgccNaattgccc
930	IGR3350a_3	t/c	T in Reference	Verified	genomic	88		gaatgtNttccatt
931	IGR3350a_3	g/c	G in reference	Verified	genomic	236		gatttgNtctctg
932	IGR3350a_4	t/c	C in reference	Verified	genomic	247		tgtttgNctgttg
933	IGR3350a_7	t/c	C in reference	Verified	genomic	321		agtgcNtatcag
934	IGR3350a_8	g/c	C in reference	verified	genomic	349		ggctgaNacaatg
935	IGR3352a_1	a/g	A in reference	Verified	genomic	282		gggatacNgTggTg
936	IGR3352a_2	t/c	T in reference	verified	genomic	314		ttattgNgTctat
937	IGR3352a_3	t/c	T in reference	Verified	genomic	330		attcttNtctctt
938	IGR3352a_4	a/g		Not yet verified	genomic	177		tgggagNgTgtat
939	IGR3352a_5	t/c	T in reference	Not yet verified	genomic	339		tcttttNttcttt
940	IGR3352a_6	a/g	G in ref	Verified	genomic	547		gtgtcaNttttgga

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
941	IGR3370a_1	other/w+	poly t	Verified	genomic	354		ctttccNtttttt
942	IGR3370a_2	t/c		Verified	genomic	378		agacagNctgctc
943	IGR3374a_1	a/t		Verified	genomic	193		ggctatNgaaaaa
944	IGR3378a_1	a/t		Verified	genomic	418		gcgggNcataaag
945	IGR3378a_2	a/t		Verified	genomic	563		cttttNaaaaatagg
946	IGR3388a_1	a/t		Not yet verified	genomic	48		aatcttaNagtacatt
947	IGR3388a_2	other/w+		Not yet verified	genomic	441		gttgaaatcNtttttttttt
948	IGR3394a_1	t/c		Not yet verified	genomic	493		ccctacaNgtaaat
949	IGR3406a_1	t/c		Not yet verified	genomic	158		aaatataNacatttat
950	IGR3420a_1	a/g		Not yet verified	genomic	322		cgccttcNataaaa
951	IGR3428a_2	a/g	A in ref	Not yet verified	genomic	278		tataaggNtaagg
952	IGR3454a_1	t/c		Not yet verified	genomic	382		tcccatNtgtaggtt
953	IGR3454a_2	t/c		Not yet verified	genomic	450		aattagaNcccattt
954	IGR3456a_1	a/g	G in Reference	Not yet verified	genomic	42		tcttcNtttggt
955	IGR3456a_2	a/t	T in Reference	Not yet verified	genomic	81		gtgggtNgtagtt
956	IGR3456a_3	t/c	T in reference	Not yet verified	genomic	108		ccttcaNgtccct
957	IGR3456a_4	a/g	G in Reference	Not yet verified	genomic	109		cttcatNtccct
958	IGR3456a_5	t/g	T in reference	Not yet verified	genomic	178		actcatNgtttgg
959	IGR3456a_6	a/g	G in Reference	Not yet verified	genomic	179		ctcattNtttggc

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
960	IGR3456a_7	other/w+	poly T	Not yet verified	genomic	204		ctctctcNtttttt
961	IGR3456a_9	a/g		Not yet verified	genomic	398		catgtcNccigcaaa
962	IGR3460a_1	g/c	C in Reference	Not yet verified	genomic	118		tttggNgcagag
963	IGR3462a_1	other/w+	poly t	Not yet verified	genomic	419		ttttttNgctcatca
964	IGR3466a_1	a/g		Not yet verified	genomic	185		gcctcatgNacccctacc
965	IGR3466a_2	g/c		Not yet verified	genomic	503		ctcacaaaNgctccagt
966	IGR3470a_1	a/t		Not yet verified	genomic	199		gctggaggNaggactag
967	IGR3470a_2	t/c		Not yet verified	genomic	438		gtcaaatNattcattt
968	IGR3470a_3	a/t		Not yet verified	genomic	178		tccagctgaNgatgcagg
969	IGR3470a_4	a/t		Not yet verified	genomic	214		agccctgaNtgggcacca
970	IGR3475a_1	a/c		Verified	genomic	483		caccctgctNtatacact
971	IGR3475a_2	t/c		Verified	genomic	490		ctatacaNtggttggt
972	IGR3477a_1	t/g		Verified	genomic	468		ccagggtcaagNtgcctgagt
973	IGR3479a_1	a/g		Verified	genomic	178		tagtgggtgNagtctgggc
974	IGR3479a_2	a/t		Not yet verified	genomic	229		cgtacccaNtagglaa
975	IGR3481a_1	a/t		Not yet verified	genomic	139		ctgggaggaNctggggact
976	IGR3483a_1	a/g		Verified	genomic	539		gtgatgggNtgcctaaag
977	IGR3483a_2	t/g		Verified	genomic	495		atctggttaaNaagggtgg
978	IGR3483a_3	a/c		Not yet verified	genomic	135		gaaacaggNgcggltggca
979	IGR3485a_1	a/g		Verified	genomic	198		ctctgaaggNtcatcacag
980	IGR3487a_1	a/t		Not yet verified	genomic	184		ctctccaagcNcctgctagta

Table 3

SEQ. ID NO.	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
981	IGR3493a_1	a/c		Not yet verified	genomic	434		ctctttattatNtccctttccca
982	IGR3493a_2	t/c		Verified	genomic	517		gatgaagggaNtgggcatgc
983	IGR3493a_3	a/g		Not yet verified	genomic	534		tgctaccaNgtgagcca
984	IGR3493a_4	g/c		Not yet verified	genomic	542		aggtagcccaNcaggatgag
985	IGR3495a_1	a/g		Not yet verified	genomic	195		tcacaacaNaagtcacg
986	IGR3499a_1	t/c		Verified	genomic	467		aggggtggactgNtctgttct
987	IGR3501a_1	other/w+	poly T	Not yet verified	genomic	503		ttcatgaagaacNttttttttttg
988	IGR3505a_1	t/c		Verified	genomic	213		tgccctctcNgggcccctgggg
989	IGR3515a_1	a/g		Verified	genomic	604		agtggtgggagNaaacctccagc
990	IGR3515a_2	t/g		Not yet verified	genomic	440		ciccttcgNccagggtga
991	IGR3519a_1	t/c		Verified	genomic	230		ttctctcctNccctgctca
992	IGR3523a_1	t/c	T in ref	Verified	genomic	193		cacggccagNagcctcttg
993	IGR3525a_1	t/g		Verified	genomic	322		ttcagaggggggNggcgggtcaagt
994	IGR3527a_1	t/c		Verified	genomic	87		cacgtggNctgagctg
995	IGR3529a_1	a/g		Verified	genomic	251		catgtacaggNgacagatctgg
996	IGR3529a_2	t/c		Verified	genomic	120		aagtgctNgaatgga
997	IGR3531a_1	t/c		Not yet verified	genomic	361		tcaacccctgNtctgtacaa
998	IGR3533a_1	a/g		Verified	genomic	137		cacagggagNgttgaga
999	IGR3535a_1	g/c		Not yet verified	genomic	462		ctgtctcNgtggggagga
1000	IGR3535a_2	a/t		Not yet verified	genomic	363		tcgagcctgNctgatggcaaa
1001	IGR3537a_1	a/g		Verified	genomic	426		ggaggtgtgagNgcagaagt
1002	IGR3551a_1	a/c		Not yet verified	genomic	403		ggccatccaNcagaaac
1003	IGR3553a_2	a/g		Verified	genomic	125		tccccacNctgatcac
1004	IGR3553a_3	t/c		Verified	genomic	426		gaattgacctaNggagtcgc
1005	IGR3555a_1	t/c		Verified	genomic	188		actgcagcctNgacctcca

Table 3

SEQ. ID NO.	Polyorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
1026	IL9_6085	c/t		Verified	gene	n/a	Interleukin 9	ttctctggaacagccatgcaaccaaaccaNggcaggcaacgcgctgaca t
1027	IL9_6085	c/t		Verified	gene	n/a	Interleukin 9	ccctggaacagccatgcaaccaaaccaNggcaggcaacgcgctgacat
1028	IRF1ex1_1	g/c		Verified	gene	n/a	Interferon regulatory factor 1	aagacgtgcgccccgagccccgcggaanCgaggccaccgccggagccgtg cccgat
1029	IRF1pro1_2	c/t		Verified	gene	n/a	Interferon regulatory factor 1	cacggggcagggtaggctttctgctNcttcactccccaggcagggtgagt
1030	IRFex6_1	t/c		Verified	gene	n/a	Interferon regulatory factor 1	ctgacctgagggtggtcctccagacct
1031	IRFex9_1	t/g		Verified	gene	n/a	Interferon regulatory factor 1	gccactcgcactnctccaagagctg
1032	IRFex9_2	t/g		Verified	gene	n/a	Interferon regulatory factor 1	tccatccacgtttttgtgctgccactc
1033	IRFpro1_1	a/g		Verified	gene	n/a	Interferon regulatory factor 1	gtagggtatattttatgtgt
1034	IRFpro1_2	t/c		Verified	gene	n/a	Interferon regulatory factor 1	ggggcagggtaggctttctgctNcttcactccccaggcagggtgagt
1035	OCTex5_1	a/g		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	gaatcaaatatcactgctgglacagctNlgtgttcatcttttgcagctttttgga
1036	OCTN1ex1_1	g/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	gctgttagaaattggggcgcggaanCccggggaccgttccctgggaaaca

Table 3

SEQ. ID NO:	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
1037	OCTN1ex3_1	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	gccctgagtcaggcatcaatgcagaNttagtgtttttcagggtcttgccag
1038	OCTN1ex6_1	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	ggatatctgcattccagggtcacttattaNttaccatagcagcaaaagacataa tgg
1039	OCTN1ex6_2	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	cttatgcatgcaactctcacttcacctgac
1040	OCTN1ex9_1	a/g		Verified	gene	n/a	carnitine transporter (organic cation transporter) 1	ctcagtcatgctgacagatgttcctttgNtagagttctttgccctaccagagttct c
1041	OCTN2ex1_1	a/g		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	gtcgcgccccggctccagccccNagcgcgcgagagttggcgatgg
1042	OCTN2ex3_1	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	accctgtcccccttgaggacatcacagNtgtctccagaagaggtaggatg g
1043	OCTN2ex3_2	a/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	ttctcggtctcacagtgcccatgcta

Table 3

SEQ. ID NO:	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on genomic reference sequence	Gene Name	Flanking Sequence
1044	OCTN2ex4_1	a/g		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	gccagtgggcacatggggcacannggtcacactcaccaccaga
1045	OCTN2ex4_2	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	actcaccaccagatgccacgcaNagcaccgccgcatcgtcagcgcc
1046	OCTN2ex6_1	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	aactccctaggccctgtcagtaaNaaatcagagtgaaatgaaatgagga
1047	OCTN2ex6_2	t/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	tatcctttcactctctgatgacaNaggctttgaatttctcagggc
1048	OCTN2ex7_1	g/c		Verified	gene	n/a	carnitine transporter (organic cation transporter) 2	gcaagttaggaglatcaagcgaaaNccaaaatagcccactgatatggc
1049	Polyex3_1	t/g		Verified	gene	n/a	Prolyl 4 hydroxylase	gcctataaggaggaaacctttgagaggNtgatgtggggctggcctggttactt catg
1050	Polyex6_1	t/c		Verified	gene	n/a	Prolyl 4 hydroxylase	ctatccagtggtcaggcttctcttgaagNgggaatctcttccctaatacca
1051	Rad50ex16_1	ins/del	ins/del caa	Verified	gene	n/a	RAD50	tcctctcgtaNaaagactgaa
1052	Rad50ex16_2	other/w+	ta repeat	Verified	gene	n/a	RAD50	agactgctcNaaaaataaaa
1053	Rad50ex16_3	ins/del	ins/del t	Verified	gene	n/a	RAD50	ttaaaataatttNacaaaaaacat
1054	Rad50ex25a_1	other/w+	ca repeat	Verified	gene	n/a	RAD50	atttaggaNccccccc
1055	Rad50ex4_1	t/c		Verified	gene	n/a	RAD50	cccttctgttttaaaNttttctgttaaaaaag

Table 3

SEQ. ID NO:	Polymorphism Name	Poly type	comment #1 Polymorphisms details	comment #2	comment #3 Verification Status	comment #4 Position on reference genomic sequence	Gene Name	Flanking Sequence
1056	Rad50ex4_2	t/c		Verified	gene	n/a	RAD50	ttaatggactacaaaagtNtattaagggttacaa
1057	Rad50ex7_1	a/t		Verified	gene	n/a	RAD50	gagattcttcattcaNacagaaaaatgtataacat
1058	Sept2ex10b_1	a/g		Verified	gene	n/a	Septin-like	ttctaaatatttatttgNcaccagcgtaagacaa
1059	Sept2ex10c_1	a/g		Verified	gene	n/a	Septin-like	attaagactcccaagcNaatcctgcataattccaa
1060	Sept2ex10d_1	t/c		Verified	gene	n/a	Septin-like	gtgtgtccacNgaggcacgg
1061	Sept2ex10f_1	a/g		Verified	gene	n/a	Septin-like	tcctgttaagtNgggctcatgga
1062	Sept2ex2_1	g/c		Not yet verified	gene	n/a	Septin-like	tgtcagggccctgNcctcagaca
1063	Sept2ex3_1	t/c		Verified	gene	n/a	Septin-like	cccagacctaNgacctccagga
1064	TCF_1625	c/t		Verified	gene	n/a	t cell transcription factor 1	cactttgcctgcaggTgcaccgaaaggacNtgggggataaaaattcaaaaa agtgat

Table 4. Summary of best SNPs in chromosome 5 region.

	SNP marker name	Approximate Physical position ¹	SNP type	Transmitted allele	Frequency of allele ²	Transmitted	Untransmitted	C ²	p-value
5	IGR2055 a 1	435.0	G/T	G	0.357	87	39	18.29	0.000019
	IGR2060 a 1	437.5	C/G	C	0.351	81	34	19.21	0.000012
10	IGR2063 b 1	439.0	C/G	G	0.359	87	37	20.16	0.000007
	IGR2069 a 2	442.0	C/T	T	0.627	52	20	14.22	0.00016
	IGR2078 a 1	446.5	A/G	A	0.364	48	16	16.00	0.000063
15	IGR2096 a 1	455.5	A/C	A	0.349	75	32	17.28	0.000032
	IGR2198 a 1	506.5	C/G	G	0.364	87	41	16.53	0.000048
20	IGR2230 a 1	522.5	C/T	T	0.415	67	28	16.01	0.000063
	IGR2277 a 1	546.0	A/G	G	0.417	79	37	15.21	0.000096
	IGR3081 a 1	609.0	G/T	G	0.338	79	35	16.98	0.000038
25	IGR3096 a 1	616.5	C/T	C	0.429	89	42	16.86	0.00004
	PROLYL ex3 1	686.5	G/T	T	0.383	79	39	13.56	0.00023

¹ Position (kb) on the 850 kb reference sequence.

30 ² Frequency of allele calculated from the untransmitted parental chromosomes.

TABLE 5

>IGR2001a SEQ ID NO: 1065

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>IGR2002a SEQ ID NO: 1066

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>IGR2003a SEQ ID NO: 1067

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>IGR2015a SEQ ID NO: 1079

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From the foregoing, it is apparent that the invention includes a number of general uses that can be expressed concisely as follows. The invention provides for the use of any of the nucleic acid molecules described herein in the diagnosis or monitoring of diseases, particularly IBD, such as in the genotyping of samples from
5 individuals to be tested. The invention further provides for the use of any of the nucleic acid molecules in the manufacture of a medicament for the treatment or prophylaxis of such diseases. The invention further provides for the use of any of the nucleic acid molecules as a pharmaceutical.

While this invention has been particularly shown and described with references
10 to preferred embodiments thereof, it will be understood by those skilled in the art that various changes in form and details may be made therein without departing from the scope of the invention encompassed by the appended claims.